```
19971027 (60)
                        US 1997-63329P
                                            19971027 (60)
                        US 1997-63327P
                                            19971028 (60)
                        US 1997-63549P
                                            19971028 (60)
                        US 1997-63541P
                                            19971028 (60)
                        US 1997-63550P
                                            19971028 (60)
                        US 1997-63542P
                                            19971028 (60)
                        US 1997-63544P
                                            19971028 (60)
                        US 1997-63564P
                                            19971029 (60)
                        US 1997-63734P
                                            19971029 (60)
                        US 1997-63738P
                                            19971029 (60)
                        US 1997-63704P
                                            19971029 (60)
                        US 1997-63435P
                                            19971029 (60)
                        US 1997-64215P
                                            19971029 (60)
                        US 1997-63735P
                                            19971029 (60)
                        US 1997-63732P
                                            19971031 (60)
                        US 1997-64103P
                        US 1997-63870P
                                            19971031 (60)
                        US 1997-64248P
                                            19971103 (60)
                                            19971107 (60)
                        US 1997-64809P
                        US 1997-65186P
                                            19971112 (60)
                                            19971117 (60)
                        US 1997-65846P
                        US 1997-65693P
                                            19971118 (60)
                        US 1997-66120P
                                            19971121 (60)
                                            19971121 (60)
                        US 1997-66364P
                                            19971124 (60)
                        US 1997-66772P
                                            19971124 (60)
                        US 1997-66466P
                                            19971124 (60)
                        US 1997-66770P
                        US 1997-66511P
                                            19971124 (60)
                        US 1997-66453P
                                            19971124 (60)
                        Utility
                        APPLICATION
                        BRINKS HOFER GILSON & LIONE, P.O. BOX 10395, CHICAGO,
                        IL, 60610
                        38
                        124 Drawing Page(s)
                        21263
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
     (FILE 'HOME' ENTERED AT 10:30:42 ON 15 JAN 2003)
     FILE 'MEDLINE, DGENE, EMBASE, SCISEARCH, USPATFULL, WPIDS, JICST-EPLUS,
     FSTA' ENTERED AT 10:31:56 ON 15 JAN 2003
           2316 S BONE MORPHOGENIC PROTEIN
          26968 S ARTICULAR CARTILAGE
            125 S OSTEOCHONDRAL GRAFT
              0 S L1 () L2 () L3
             76 S L1 AND L2
              1 S L5 AND L3
              1 S L3 AND L1
=> s 13 and regeneration
             4 L3 AND REGENERATION
=> d 18 ti abs ibib tot
     ANSWER 1 OF 4 USPATFULL
       Device for regeneration of articular cartilage and other
       An implantable device for facilitating the healing of voids in bone, `
       cartilage and soft tissue is disclosed. A preferred embodiment includes
```

DOCUMENT TYPE:

LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS:

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

LINE COUNT:

=> d his

L1

L2L3

L4L5

L6

**L7** 

L8

ΤI

AB

FILE SEGMENT:

a cartilage region comprising a polyelectrolytic complex joined with a subchondral bone region. The cartilage region, of this embodiment, enhances the environment for chondrocytes to grow articular cartilage; while the subchondral bone region enhances the environment for cells which migrate into that region's macrostructure and which differentiate into osteoblasts. A hydrophobic barrier exists between said regions, of this embodiment. In one embodiment, the polyelectrolytic complex transforms to hydrogel, following the implant procedure.

ACCESSION NUMBER:

2002:55324 USPATFULL

TITLE:

Device for regeneration of articular

cartilage and other tissue

INVENTOR(S):

Brekke, John H., Duluth, MN, UNITED STATES Goldman, Scott M., Paoli, PA, UNITED STATES

NUMBER KIND DATE -----PATENT INFORMATION:

APPLICATION INFO .:

US 2002032488 A1 20020314 US 2001-909027 A1 20010719 (9)

RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 1998-206604, filed on 7 Dec 1998, GRANTED, Pat. No. US 6264701 Division of

Ser. No. US 1994-242557, filed on 13 May 1994, GRANTED,

Pat. No. US 5981825

DOCUMENT TYPE: FILE SEGMENT:

Utility APPLICATION

LEGAL REPRESENTATIVE:

Alan D. Kamrath, Kensey Nash Corporation, 55 E. Uwchlan

Avenue, Exton, PA, 19341

NUMBER OF CLAIMS:

1349

EXEMPLARY CLAIM: LINE COUNT:

ANSWER 2 OF 4 USPATFULL L8

Scaffold matrix and tissue maintaining systems TI

AB The invention concerns a scaffold which is used as a growth supportive base for various cells and tissue explants from three-dimensional tissue comprising naturally derived connective or skeletal tissue into attached flakes having a very high porosity. Alternatively the scaffold is

composed of fused epiphyses.

ACCESSION NUMBER:

2002:16925 USPATFULL

TITLE:

Scaffold matrix and tissue maintaining systems

INVENTOR(S):

Nevo, Zvi, Herzliya, ISRAEL Robinson, Dror, Shimshon, ISRAEL

PATENT ASSIGNEE(S):

RAMOT UNIVERSITY AUTHORITY FOR APPLIED RESEARCH & INDUSTRIAL DEVELOPMENT LTD. (non-U.S. corporation)

NUMBER KIND DATE \_\_\_\_\_\_ US 2002009805 A1 20020124 US 2001-826389 A1 20010404 20010404 (9)

APPLICATION INFO.: RELATED APPLN. INFO.:

PATENT INFORMATION:

Continuation-in-part of Ser. No. US 1999-345138, filed

on 6 Jul 1999, PENDING

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

LEGAL REPRESENTATIVE:

LADAS & PARRY, 26 WEST 61ST STREET, NEW YORK, NY, 10023

NUMBER OF CLAIMS: 32 EXEMPLARY CLAIM:

10 Drawing Page(s)

NUMBER OF DRAWINGS: LINE COUNT:

903

L8 ANSWER 3 OF 4 USPATFULL

Multi-stage collagen-based template or implant for use in the repair of TI cartilage lesions

The invention is a template to aid in the regeneration of AB

articular cartilage. The template is formed by combining a porous collagen sponge ("collagen matrix") with a dense collagen membrane. The dense collagen membrane is placed on the surface of the cartilage defect to prevent cell migration from the subchondral plate and vasculature. The collagen membrane will allow movement and exchange of fluids, nutrients, cytokines and other factors necessary for cartilage regeneration. The collagen matrix has been developed to allow attachment and growth of cells, specifically chondrocytes which are normally found in articular cartilage. The collagen matrix can be combined with chondrocytes in vitro, and therefore serve to transport cultured cells to the defect site and to retain the cells in position following implantation. Procedures are described to effectively use the two-staged template, and to fix the template to the repair site.

ACCESSION NUMBER:

2000:80202 USPATFULL

TITLE:

Multi-stage collagen-based template or implant for use

in the repair of cartilage lesions

INVENTOR(S):

Pachence, James M., Hopewell, NJ, United States

Frenkel, Sally, Flushing, NY, United States Menche, David, New York, NY, United States

PATENT ASSIGNEE(S):

The Hospital for Joint Disease Orthopaedic Institute,

New York, NY, United States (U.S. corporation)

NUMBER	KIND	DATE

PATENT INFORMATION:

US 6080194 20000627 US 1995-385290

APPLICATION INFO.: DOCUMENT TYPE: Utility 19950210 (8)

FILE SEGMENT: Granted

PRIMARY EXAMINER:

Prebilic, Paul B.

LEGAL REPRESENTATIVE:

Caesar, Rivise, Bernstein, Cohen & Pokotilow, Ltd.

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

AB

4 Drawing Figure(s); 4 Drawing Page(s)

LINE COUNT:

ANSWER 4 OF 4 JICST-EPlus COPYRIGHT 2003 JST

636

- Four Case Reports of Mosaicplasty for Knee Joint. TI
  - Repairing a defect or injury of articular cartilage is a significant challenge. Osteochondral graft, periosteal transplantation, drilling, and chondrocyte transplantation have been attempted clinically for articular surface defects. We evaluated repairs of articular cartilage by mosaicplasty. Four knees of 4 patients (2 men and 2 women) that underwent mosaicplasty were evaluated in this series. Mean patient age at surgery was 41 years. All knees underwent follow-up MRI, 2 knees underwent follow-up arthroscopy and needle biopsy after informed consent was obtained. The mean period from surgery to final follow-up was 21 months. The mean period from surgery to follow-up arthroscopy was 11 months. Four cases of mosaicplasty presented satisfactory regeneration of the articular cartilage as seem by MRI or arthroscopic examination. Two knees, after receiving mosaicplasty, demonstrated regeneration of hyaline cartilage even around the gaps in mosaicplasty, by needle biopsy. However, the structure of hyaline cartilage around the gaps in mosaicplasty differed from that of normal hyaline cartilage. Several reports described a good clinical outcome of mosaicplasty. However, only Hangody reported good hyaline cartilage regeneration at the recipient site and fibrous cartilage at the donor site. Our results demonstrated regeneration of the hyaline cartilage in the gap area of mosaicplasty, but the structure of hyaline cartilage differed from normal. There is a risk of renewed degeneration due to the poor structure of hyaline cartilage. Mosaicplasty is a sure method of repairing hyaline cartilage where there is a small defect in the articular surface. However, one report pointed out the risk of articular degeneration at the donor site after mosaicplasty. One of our cases

demonstrated bony defect at the donor site 21 months after mosaicplasty. Adequate observation of both the donor site and recipient site is needed

after mosaicplasty. (author abst.) ACCESSION NUMBER: 1010895536 JICST-EPlus

Four Case Reports of Mosaicplasty for Knee Joint. TITLE:

ICHINOHE SADAFUMI; KOYAMA AKIKO; ENDO TAKESHI; KITAGAWA **AUTHOR:** 

YUKA; YOSHIDA MASAAKI; SHIMAMURA TADASHI

SHIROKURA YOSHIHIRO; HONDA KEI

Iwateidai Seikeigeka CORPORATE SOURCE:

Moriokashibyoin Seikeigeka

Nippon Riumachi, Kansetsu Geka Gakkai Zasshi (Japanese SOURCE:

> Journal of Rheumatism and Joint Surgery), (2001) vol. 20, no. 2, pp. 169-175. Journal Code: Y0692A (Fig. 6, Ref. 10)

ISSN: 0287-3214

PUB. COUNTRY: Japan

Journal; Short Communication DOCUMENT TYPE:

LANGUAGE: Japanese STATUS: New

=> d his

(FILE 'HOME' ENTERED AT 10:30:42 ON 15 JAN 2003)

FILE 'MEDLINE, DGENE, EMBASE, SCISEARCH, USPATFULL, WPIDS, JICST-EPLUS,

FSTA' ENTERED AT 10:31:56 ON 15 JAN 2003

2316 S BONE MORPHOGENIC PROTEIN L1

L226968 S ARTICULAR CARTILAGE

125 S OSTEOCHONDRAL GRAFT L3

0 S L1 () L2 () L3 L4

76 S L1 AND L2  $L_5$ 

1 S L5 AND L3 1.6

1 S L3 AND L1 L7

4 S L3 AND REGENERATION

=> s 15 and regeneration

L9 44 L5 AND REGENERATION

=> d 19 ti abs ibib 1-15

ANSWER 1 OF 44 MEDLINE L9

Cartilage and bone regeneration using gene-enhanced tissue TТ

engineering.

Joint cartilage injury remains a major problem in orthopaedics with more AB than 500,000 cartilage repair procedures performed yearly in the United States at a cost of hundreds of millions of dollars. No consistently

reliable means to regenerate joint cartilage currently exists. The technologies of gene therapy and tissue engineering were combined using a

retroviral vector to stably introduce the human bone morphogenic protein-7 complementary deoxyribonucleic

acid into periosteal-derived rabbit mesenchymal stem cells. Bone

morphogenic protein-7 secreting gene modified cells

subsequently were expanded in monolayer culture, seeded onto polyglycolic acid grafts, implanted into a rabbit knee osteochondral defect model, and evaluated for bone and cartilage repair after 4, 8, and 12 weeks. The

grafts containing bone morphogenic protein-7

gene modified cells consistently showed complete or near complete bone and articular cartilage regeneration at 8 and 12

weeks whereas the grafts from the control groups had poor repair as judged by macroscopic, histologic, and immunohistologic criteria. This is the first report of articular cartilage

regeneration using a combined gene therapy and tissue engineering

approach. MEDLINE ACCESSION NUMBER: 2000488818

PubMed ID: 11039767 DOCUMENT NUMBER: 20492911

Cartilage and bone regeneration using TITLE:

gene-enhanced tissue engineering.

Mason J M; Breitbart A S; Barcia M; Porti D; Pergolizzi R AUTHOR:

G; Grande D A

Department of Research, North Shore University Hospital-New CORPORATE SOURCE:

York University School of Medicine, Manhasset 11030, USA.

CLINICAL ORTHOPAEDICS AND RELATED RESEARCH, (2000 Oct) (379 SOURCE:

Suppl) S171-8.

Journal code: 0075674. ISSN: 0009-921X.

United States PUB. COUNTRY:

Journal; Article; (JOURNAL ARTICLE) DOCUMENT TYPE:

LANGUAGE: English

Abridged Index Medicus Journals; Priority Journals FILE SEGMENT:

ENTRY MONTH: 200011

Entered STN: 20010322 ENTRY DATE:

> Last Updated on STN: 20010322 Entered Medline: 20001103

ANSWER 2 OF 44 EMBASE COPYRIGHT 2003 ELSEVIER SCI. B.V. L9

Cartilage and bone regeneration using gene-enhanced tissue ΤI

engineering.

Joint cartilage injury remains a major problem in orthopaedics with more AB than 500,000 cartilage repair procedures performed yearly in the United States at a cost of hundreds of millions of dollars. No consistently reliable means to regenerate joint cartilage currently exists. The technologies of gene therapy and tissue engineering were combined using a retroviral vector to stably introduce the human bone

morphogenic protein-7 complementary deoxyribonucleic

acid into periosteal-derived rabbit mesenchymal stem cells. Bone

morphogenic protein-7 secreting gene modified cells

subsequently were expanded in monolayer culture, seeded onto polyglycolic acid grafts, implanted into a rabbit knee osteochondral defect model, and evaluated for bone and cartilage repair after 4, 8, and 12 weeks. The grafts containing bone morphogenic protein-7

gene modified cells consistently showed complete or near complete bone and articular cartilage regeneration at 8 and 12

weeks whereas the grafts from the control groups had poor repair as judged by macroscopic, histologic, and immunohistologic criteria. This is the first report of articular cartilage

regeneration using a combined gene therapy and tissue engineering approach.

2000362559 EMBASE ACCESSION NUMBER:

TITLE: Cartilage and bone regeneration using

gene-enhanced tissue engineering.

Mason J.M.; Breitbart A.S.; Barcia M.; Porti D.; Pergolizzi AUTHOR:

R.G.; Grande D.A.

Dr. J.M. Mason, Gene Therapy Vector Laboratory, Department CORPORATE SOURCE:

of Research, North Shore University Hospital, 350 Community

Drive, Manhasset, NY 11030, United States

Clinical Orthopaedics and Related Research, (2000) -/379 SOURCE :

SUPPL. (S171-S178).

Refs: 27

ISSN: 0009-921X CODEN: CORTBR

COUNTRY: United States

DOCUMENT TYPE: Journal; Conference Article FILE SEGMENT: 022 Human Genetics 033 Orthopedic Surgery

> 036 Health Policy, Economics and Management

LANGUAGE: English SUMMARY LANGUAGE: English

1.9 ANSWER 3 OF 44 SCISEARCH COPYRIGHT 2003 ISI (R)

ΤI Cartilage and bone regeneration using gene-enhanced tissue engineering

AΒ

Joint cartilage injury remains a major problem in orthopaedics with more than 500,000 cartilage repair procedures performed yearly in the United States at a cost of hundreds of millions of dollars. No consistently reliable means to regenerate joint cartilage currently exists. The technologies of gene therapy and tissue engineering were combined using a retroviral vector to stably introduce the human hone morphogenic protein-7 complementary deoxyribonucleic acid into periosteal-derived rabbit mesenchymal stem cells. Bone

morphogenic protein-7 secreting gene modified cells subsequently were expanded in monolayer culture, seeded onto polyglycolic acid grafts, implanted into a rabbit knee osteochondral defect model, and evaluated for bone and cartilage repair after 4, 8, and 12 weeks, The grafts containing bone morphogenic protein-7

gene modified cells consistently showed complete or near complete bone and articular cartilage regeneration at 8 and 12

weeks whereas the grafts from the control groups had poor repair as judged by macroscopic, histologic, and immunohistologic criteria, This is the first report of articular cartilage

regeneration using a combined gene therapy and tissue engineering approach.

ACCESSION NUMBER: 2000:777821 SCISEARCH

THE GENUINE ARTICLE: 362NP

TITLE: Cartilage and bone regeneration using

gene-enhanced tissue engineering

AUTHOR: Mason J M (Reprint); Breitbart A S; Barcia M; Porti D;

Pergolizzi R G; Grande D A

CORPORATE SOURCE: NYU, GENE THERAPY VECTOR LAB, DEPT RES, N SHORE UNIV HOSP,

SCH MED, 350 COMMUNITY DR, MANHASSET, NY 11030 (Reprint); NYU, DIV PLAST & RECONSTRUCT SURG, SCH MED, N SHORE UNIV HOSP, MANHASSET, NY 11030; NYU, DIV ORTHOPED SURG, SCH MED, N SHORE UNIV HOSP, DEPT SURG, MANHASSET, NY 11030

COUNTRY OF AUTHOR:  $\lambda$ 

SOURCE:

CLINICAL ORTHOPAEDICS AND RELATED RESEARCH, (OCT 2000) No.

379, Supp. [S], pp. S171-S178.

Publisher: LIPPINCOTT WILLIAMS & WILKINS, 530 WALNUT ST,

PHILADELPHIA, PA 19106-3621.

ISSN: 0009-921X.

DOCUMENT TYPE: Article; Journal FILE SEGMENT: LIFE; CLIN

LANGUAGE: English
REFERENCE COUNT: 27

\*ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS\*

L9 ANSWER 4 OF 44 USPATFULL

TI Secreted and transmembrane polypeptides and nucleic acids encoding the

The present invention is directed to novel polypeptides and to nucleic acid molecules encoding those polypeptides. Also provided herein are vectors and host cells comprising those nucleic acid sequences, chimeric polypeptide molecules comprising the polypeptides of the present invention fused to heterologous polypeptide sequences, antibodies which bind to the polypeptides of the present invention and to methods for producing the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT. ACCESSION NUMBER: 2003:3495 USPATFULL

TITLE: Secreted and transmembrane polypeptides and nucleic

acids encoding the same

INVENTOR(S): Ashkenazi, Avi, San Mateo, CA, UNITED STATES

Botstein, David, Belmont, CA, UNITED STATES
Desnoyers, Luc, San Francisco, CA, UNITED STATES
Eaton, Dan L., San Rafael, CA, UNITED STATES

Ferrara, Napoleone, San Francisco, CA, UNITED STATES

Filvaroff, Ellen, San Francisco, CA, UNITED STATES Fong, Sherman, Alameda, CA, UNITED STATES Gao, Wei-Qiang, Palo Alto, CA, UNITED STATES Gerber, Hanspeter, San Francisco, CA, UNITED STATES Gerritsen, Mary E., San Mateo, CA, UNITED STATES Goddard, Audrey, San Francisco, CA, UNITED STATES Godowski, Paul J., Burlingame, CA, UNITED STATES Grimaldi, J. Christopher, San Francisco, CA, UNITED STATES

Gurney, Austin L., Belmont, CA, UNITED STATES
Hillan, Kenneth J., San Francisco, CA, UNITED STATES
Kljavin, Ivar J., Lafayette, CA, UNITED STATES
Mather, Jennie P., Millbrae, CA, UNITED STATES
Pan, James, Belmont, CA, UNITED STATES
Paoni, Nicholas F., Belmont, CA, UNITED STATES
Roy, Margaret Ann, San Francisco, CA, UNITED STATES
Stewart, Timothy A., San Francisco, CA, UNITED STATES
Tumas, Daniel, Orinda, CA, UNITED STATES
Williams, P. Mickey, Half Moon Bay, CA, UNITED STATES
Wood, William I., Hillsborough, CA, UNITED STATES
Genentech, Inc. (U.S. corporation)

PATENT ASSIGNEE(S):

NUMBER KIND DATE
----US 2003003530 A1 20030102
US 2001-904011 A1 20010711 (9)

NUMBER

PATENT INFORMATION: APPLICATION INFO.: RELATED APPLN. INFO.:

Continuation of Ser. No. US 2000-665350, filed on 18 Sep 2000, PENDING

DATE

# PRIORITY INFORMATION:

WO	1998-US18824	19980910	
WO	1998-US19177	19980914	
WO	1998-US19330	19980916	
WO	1998-US19437	19980917	
WO		19981201	
WO		19990908	
WO	1999-US20944	19990913	
WO			
WO			
WO			
WO		19991129	
WO		19991130	
	1999-US28301		
	1999-US28564		
	1999-US28565		
	1999-US30095		
		19991220	
	1999-US30911		
		20000105	
WO			
WO			
WO		20000224	
WO		20000302	
WO		20000320	
WO			
WO			
WO			
WO		20000728	
WO		20000824	
US		19970917	
US			
US			
US	1997-59117P	19970917	(60)

```
19970917 (60)
US 1997-59113P
                   19970917 (60)
US 1997-59121P
                   19970917 (60)
US 1997-59119P
US 1997-59263P
                   19970918 (60)
US 1997-59266P
                   19970918 (60)
US 1997-62125P
                   19971015 (60)
US 1997-62287P
                   19971017 (60)
                   19971017 (60)
US 1997-62285P
US 1997-63486P
                   19971021 (60)
US 1997-62816P
                   19971024 (60)
                   19971024 (60)
US 1997-62814P
                   19971024 (60)
US 1997-63127P
                   19971024 (60)
US 1997-63120P
                   19971024 (60)
US 1997-63121P
                   19971024 (60)
US 1997-63045P
                   19971024 (60)
US 1997-63128P
US 1997-63329P
                   19971027 (60)
                   19971027 (60)
US 1997-63327P
                   19971028 (60)
US 1997-63549P
US 1997-63541P
                   19971028 (60)
US 1997-63550P
                   19971028 (60)
US 1997-63542P
                   19971028 (60)
US 1997-63544P
                   19971028 (60)
US 1997-63564P
                   19971028 (60)
US 1997-63734P
                   19971029 (60)
US 1997-63738P
                   19971029 (60)
US 1997-63704P
                   19971029 (60)
US 1997-63435P
                   19971029 (60)
US 1997-64215P
                   19971029 (60)
US 1997-63735P
                   19971029 (60)
US 1997-63732P
                   19971029 (60)
US 1997-64103P
                   19971031 (60)
US 1997-63870P
                   19971031 (60)
US 1997-64248P
                   19971103 (60)
                   19971107 (60)
US 1997-64809P
US 1997-65186P
                   19971112 (60)
US 1997-65846P
                   19971117 (60)
                   19971118 (60)
US 1997-65693P
                   19971121 (60)
US 1997-66120P
US 1997-66364P
                   19971121 (60)
                   19971124 (60)
US 1997-66772P
US 1997-66466P
                   19971124 (60)
US 1997-66770P
                   19971124 (60)
                   19971124 (60)
US 1997-66511P
US 1997-66453P
                   19971124 (60)
```

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: BRINKS HOFER GILSON & LIONE, P.O. BOX 10395, CHICAGO,

IL, 60610

NUMBER OF CLAIMS: 38 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 124 Drawing Page(s)

LINE COUNT: 21255

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

#### L9 ANSWER 5 OF 44 USPATFULL

TI Secreted and transmembrane polypeptides and nucleic acids encoding the same

The present invention is directed to novel polypeptides and to nucleic acid molecules encoding those polypeptides. Also provided herein are vectors and host cells comprising those nucleic acid sequences, chimeric polypeptide molecules comprising the polypeptides of the present invention fused to heterologous polypeptide sequences, antibodies which bind to the polypeptides of the present invention and to methods for

producing the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2002:344632 USPATFULL

TITLE: Secreted and transmembrane polypeptides and nucleic

acids encoding the same

INVENTOR(S): Ashkenazi, Avi, San Mateo, CA, UNITED STATES

Botstein, David, Belmont, CA, UNITED STATES Desnoyers, Luc, San Francisco, CA, UNITED STATES

Eaton, Dan L., San Rafael, CA, UNITED STATES

Ferrara, Napoleone, San Francisco, CA, UNITED STATES Filvaroff, Ellen, San Francisco, CA, UNITED STATES

Fong, Sherman, Alameda, CA, UNITED STATES

Gao, Wei-Qiang, Palo Alto, CA, UNITED STATES

Gerber, Hanspeter, San Francisco, CA, UNITED STATES Gerritsen, Mary E., San Mateo, CA, UNITED STATES Goddard, Audrey, San Francisco, CA, UNITED STATES Godowski, Paul J., Burlingame, CA, UNITED STATES Grimaldi, J. Christopher, San Francisco, CA, UNITED

STATES

Gurney, Austin L., Belmont, CA, UNITED STATES

Hillan, Kenneth J., San Francisco, CA, UNITED STATES

Kljavin, Ivar J., Lafayette, CA, UNITED STATES Mather, Jennie P., Millbrae, CA, UNITED STATES

Pan, James, Belmont, CA, UNITED STATES

Paoni, Nicholas F., Belmont, CA, UNITED STATES Roy, Margaret Ann, San Francisco, CA, UNITED STATES Stewart, Timothy A., San Francisco, CA, UNITED STATES

Tumas, Daniel, Orinda, CA, UNITED STATES

Williams, P. Mickey, Half Moon Bay, CA, UNITED STATES Wood, William I., Hillsborough, CA, UNITED STATES

Genentech, Inc. (U.S. corporation)

PATENT ASSIGNEE(S):

NUMBER KIND DATE

PATENT INFORMATION: APPLICATION INFO.:

US 2002198366 A1 20021226 US 2001-907841 A1 20010717 (9)

RELATED APPLN. INFO.:

Continuation of Ser. No. US 2000-665350, filed on 18

חאתב

Sep 2000, PENDING

MIIMPED

	HOLIDBIC	2.1.2
_		

# PRIORITY INFORMATION:

WO	1998-US18824	19980910
WO	1998-US19177	19980914
WO	1998-US19330	19980916
WO	1998-US19437	19980917
WO	1998-US25108	19981201
WO	1999-US20594	19990908
WO	1999-US20944	19990913
WO	1999-US21090	19990915
WO	1999-US21547	19990915
WO	1999-US23089	19991005
WO	1999-US28214	19991129
WO	1999-US28313	19991130
WO	1999-US28301	19991201
WO	1999-US28564	19991202
WO	1999-US28565	19991202
WO	1999-US30095	19991216
WO	1999-US30999	19991220
WO	1999-US30911	19991220
WO	2000-US219	20000105
WO	2000-US3565	20000211
WO	2000-US4414	20000222
WO	2000-US5004	20000224

```
WO 2000-US5841
                   20000402
                   20000320
WO 2000-US7377
WO 2000-US8439
                   20000330
WO 2000-US14042
                   20000522
WO 2000-US15264
                   20000602
WO 2000-US20710
                   20000728
WO 2000-US23328
                   20000824
US 1997-59115P
                   19970917 (60)
US 1997-59184P
                   19970917 (60)
                   19970917 (60)
US 1997-59122P
                   19970917 (60)
US 1997-59117P
                   19970917 (60)
US 1997-59113P
                   19970917 (60)
US 1997-59121P
                   19970917 (60)
US 1997-59119P
                   19970918 (60)
US 1997-59263P
US 1997-59266P
                   19970918 (60)
US 1997-62125P
                   19971015 (60)
                   19971017 (60)
US 1997-62287P
                   19971017 (60)
US 1997-62285P
                   19971021 (60)
US 1997-63486P
US 1997-62816P
                   19971024 (60)
US 1997-62814P
                   19971024 (60)
US 1997-63127P
                   19971024 (60)
US 1997-63120P
                   19971024 (60)
US 1997-63121P
                   19971024 (60)
US 1997-63045P
                   19971024 (60)
US 1997-63128P
                   19971024 (60)
US 1997-63329P
                   19971027 (60)
US 1997-63327P
                   19971027 (60)
US 1997-63549P
                   19971028 (60)
                   19971028 (60)
US 1997-63541P
US 1997-63550P
                   19971028 (60)
US 1997-63542P
                   19971028 (60)
US 1997-63544P
                   19971028 (60)
US 1997-63564P
                   19971028 (60)
US 1997-63734P
                   19971029 (60)
US 1997-63738P
                   19971029 (60)
US 1997-63704P
                   19971029 (60)
US 1997-63435P
                   19971029 (60)
US 1997-64215P
                   19971029 (60)
US 1997-63735P
                   19971029 (60)
US 1997-63732P
                   19971029 (60)
US 1997-64103P
                   19971031 (60)
US 1997-63870P
                   19971031 (60)
US 1997-64248P
                   19971103 (60)
                   19971107 (60)
US 1997-64809P
                   19971112 (60)
US 1997-65186P
                   19971117 (60)
US 1997-65846P
US 1997-65693P
                   19971118 (60)
US 1997-66120P
                   19971121 (60)
US 1997-66364P
                   19971121 (60)
US 1997-66772P
                   19971124 (60)
US 1997-66466P
                   19971124 (60)
US 1997-66770P
                   19971124 (60)
US 1997-66511P
                   19971124 (60)
US 1997-66453P
                   19971124 (60)
Utility
APPLICATION
```

```
DOCUMENT TYPE: FILE SEGMENT:
```

LEGAL REPRESENTATIVE: BRINKS HOFER GILSON & LIONE, P.O. BOX 10395, CHICAGO,

IL, 60610

NUMBER OF CLAIMS: 38 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 124 Drawing Page(s)

LINE COUNT:

21263

#### CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L9 ANSWER 6 OF 44 USPATFULL

TI Secreted and transmembrane polypeptides and nucleic acids encoding the same

The present invention is directed to novel polypeptides and to nucleic acid molecules encoding those polypeptides. Also provided herein are vectors and host cells comprising those nucleic acid sequences, chimeric polypeptide molecules comprising the polypeptides of the present invention fused to heterologous polypeptide sequences, antibodies which bind to the polypeptides of the present invention and to methods for producing the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2002:343945 USPATFULL

TITLE: Secreted and transmembrane polypeptides and nucleic

acids encoding the same

INVENTOR(S): Ashkenazi, Avi, San Mateo, CA, UNITED STATES

Botstein, David, Belmont, CA, UNITED STATES
Desnoyers, Luc, San Francisco, CA, UNITED STATES

Ferrara, Napoleone, San Francisco, CA, UNITED STATES

Filvaroff, Ellen, San Francisco, CA, UNITED STATES

Eaton, Dan L., San Rafael, CA, UNITED STATES

Fong, Sherman, Alameda, CA, UNITED STATES Gao, Wei-Qiang, Palo Alto, CA, UNITED STATES

Gerber, Hanspeter, San Francisco, CA, UNITED STATES Gerritsen, Mary E., San Mateo, CA, UNITED STATES Goddard, Audrey, San Francisco, CA, UNITED STATES Godowski, Paul J., Burlingame, CA, UNITED STATES Grimaldi, J. Christopher, San Francisco, CA, UNITED

STATES

Gurney, Austin L., Belmont, CA, UNITED STATES

Hillan, Kenneth J., San Francisco, CA, UNITED STATES

Kljavin, Ivar J., Lafayette, CA, UNITED STATES Mather, Jennie P., Millbrae, CA, UNITED STATES

Pan, James, Belmont, CA, UNITED STATES

Paoni, Nicholas F., Belmont, CA, UNITED STATES Roy, Margaret Ann, San Francisco, CA, UNITED STATES Stewart, Timothy A., San Francisco, CA, UNITED STATES

Tumas, Daniel, Orinda, CA, UNITED STATES

Williams, P. Mickey, Half Moon Bay, CA, UNITED STATES Wood, William I., Hillsborough, CA, UNITED STATES

Genentech, Inc. (U.S. corporation)

PATENT ASSIGNEE(S): Genentech, Inc.

NUMBER KIND DATE
-----US 2002197671 A1 20021226
US 2001-907824 A1 20010717 (9)

RELATED APPLN. INFO.:

PATENT INFORMATION: APPLICATION INFO.:

Continuation of Ser. No. US 2000-665350, filed on 18

19991005

19991129

Sep 2000, PENDING

WO 1999-US23089

WO 1999-US28214

		NUMBER	DATE
INFORMATION:	WO	1998-US18824	19980910
	WO	1998-US19177	19980914
	WO	1998-US19330	19980916
	WO	1998-US19437	19980917
	WO	1998-US25108	19981201
	WO	1999-US20594	19990908
	WO	1999-US20944	19990913
	WO	1999-US21090	19990915
	WO	1999-US21547	19990915
	INFORMATION:	WO WO WO WO WO WO	

```
WO 1999-US28313
                    19991130
WO 1999-US28301
                    19991201
                    19991202
WO 1999-US28564
                    19991202
WO 1999-US28565
WO 1999-US30095
                    19991216
                    19991220
WO 1999-US30999
WO 1999-US30911
                    19991220
WO 2000-US219
                    20000105
WO 2000-US3565
                    20000211
WO 2000-US4414
                    20000222
                    20000224
WO 2000-US5004
WO 2000-US5841
                    20000302
WO 2000-US7377
                    20000320
WO 2000-US8439
                    20000330
WO 2000-US14042
                    20000522
WO 2000-US15264
                    20000602
WO 2000-US20710
                    20000728
WO 2000-US23328
                    20000824
US 1997-59115P
                    19970917
                              (60)
US 1997-59184P
                    19970917
                              (60)
US 1997-59122P
                    19970917
                              (60)
US 1997-59117P
                    19970917
                              (60)
US 1997-59113P
                    19970917
                              (60)
US 1997-59121P
                    19970917
                              (60)
US 1997-59119P
                    19970917
                              (60)
US 1997-59263P
                    19970918
                              (60)
US 1997-59266P
                    19970918
                              (60)
US 1997-62125P
                    19971015
                              (60)
US 1997-62287P
                    19971017
                              (60)
US 1997-62285P
                    19971017
                              (60)
US 1997-63486P
                    19971021
                              (60)
US 1997-62816P
                    19971024
                              (60)
US 1997-62814P
                    19971024
                              (60)
US 1997-63127P
                    19971024
                              (60)
US 1997-63120P
                    19971024
                              (60)
US 1997-63121P
                    19971024
                              (60)
US 1997-63045P
                    19971024
                              (60)
                    19971024
                              (60)
US 1997-63128P
                    19971027
US 1997-63329P
                              (60)
US 1997-63327P
                    19971027
                              (60)
US 1997-63549P
                    19971028
                              (60)
US 1997-63541P
                    19971028
                              (60)
US 1997-63550P
                    19971028
                              (60)
                    19971028
US 1997-63542P
                              (60)
US 1997-63544P
                    19971028
                              (60)
                    19971028
                              (60)
US 1997-63564P
                    19971029
                              (60)
US 1997-63734P
                    19971029
                              (60)
US 1997-63738P
US 1997-63704P
                    19971029
                              (60)
                    19971029
US 1997-63435P
                              (60)
US 1997-64215P
                    19971029
                              (60)
US 1997-63735P
                    19971029
                              (60)
US 1997-63732P
                    19971029
                              (60)
                    19971031
US 1997-64103P
                              (60)
                    19971031
US 1997-63870P
                              (60)
US 1997-64248P
                    19971103
                              (60)
US 1997-64809P
                    19971107
                              (60)
US 1997-65186P
                    19971112
                              (60)
US 1997-65846P
                    19971117
                              (60)
                    19971118
                              (60)
US 1997-65693P
US 1997-66120P
                    19971121
                              (60)
US 1997-66364P
                    19971121
                              (60)
US 1997-66772P
                    19971124
                              (60)
US 1997-66466P
                    19971124 (60)
```

US 1997-66770P 19971124 (60) US 1997-66511P 19971124 (60) US 1997-66453P 19971124 (60)

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

BRINKS HOFER GILSON & LIONE, P.O. BOX 10395, CHICAGO, LEGAL REPRESENTATIVE:

IL, 60610

NUMBER OF CLAIMS: 38 EXEMPLARY CLAIM:

124 Drawing Page(s) NUMBER OF DRAWINGS:

LINE COUNT: 22162

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 7 OF 44 USPATFULL

Methods of using bone morphogenic proteins as biomarkers for determining TΙ

cartilage degeneration and aging

Methods are provided for determining cartilage degeneration, AB regeneration, or aging in a joint tissue in a patient by measuring levels of osteogenic protein-1 (OP-1) protein and/or mRNA in

synovial fluid or joint tissue. The methods according to the invention are useful for detecting, diagnosing, predicting, determining a predisposition for, or monitoring joint tissue degeneration, regeneration, or aging in a patient including inflammatory joint

disease or age-related disorders.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

2002:337321 USPATFULL ACCESSION NUMBER:

Methods of using bone morphogenic proteins as TITLE:

biomarkers for determining cartilage degeneration and

aging

Chubinskaya, Susanna, Vernon Hills, IL, UNITED STATES INVENTOR(S):

Rueger, David C., Southborough, MA, UNITED STATES Kuettner, Klaus E., Chicago, IL, UNITED STATES

NUMBER KIND DATE -----US 2002192679 A1 US 2002-81163 A1 PATENT INFORMATION: 20021219

20020220 (10) APPLICATION INFO.:

> NUMBER DATE \_\_\_\_\_

US 2001-348111P 20011109 (60) US 2001-270528P 20010221 (60) PRIORITY INFORMATION:

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

TESTA, HURWITZ & THIBEAULT, LLP, HIGH STREET TOWER, 125 LEGAL REPRESENTATIVE:

HIGH STREET, BOSTON, MA, 02110

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 15 Drawing Page(s)

LINE COUNT: 1482

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 8 OF 44 USPATFULL

Secreted and transmembrane polypeptides and nucleic acids encoding the ТT

The present invention is directed to novel polypeptides and to nucleic AB acid molecules encoding those polypeptides. Also provided herein are vectors and host cells comprising those nucleic acid sequences, chimeric polypeptide molecules comprising the polypeptides of the present invention fused to heterologous polypeptide sequences, antibodies which bind to the polypeptides of the present invention and to methods for producing the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

2002:337301 USPATFULL ACCESSION NUMBER:

Secreted and transmembrane polypeptides and nucleic TITLE:

acids encoding the same

Ashkenazi, Avi, San Mateo, CA, UNITED STATES INVENTOR (S):

Botstein, David, Belmont, CA, UNITED STATES Desnoyers, Luc, San Francisco, CA, UNITED STATES Eaton, Dan L., San Rafael, CA, UNITED STATES

Ferrara, Napoleone, San Francisco, CA, UNITED STATES Filvaroff, Ellen, San Francisco, CA, UNITED STATES

Fong, Sherman, Alameda, CA, UNITED STATES

Gao, Wei-Qiang, Palo Alto, CA, UNITED STATES

Gerber, Hanspeter, San Francisco, CA, UNITED STATES Gerritsen, Mary E., San Mateo, CA, UNITED STATES Goddard, Audrey, San Francisco, CA, UNITED STATES Godowski, Paul J., Burlingame, CA, UNITED STATES Grimaldi, J. Christopher, San Francisco, CA, UNITED

STATES

Gurney, Austin L., Belmont, CA, UNITED STATES

Hillan, Kenneth J., San Francisco, CA, UNITED STATES Kljavin, Ivar J., Lafayette, CA, UNITED STATES

Mather, Jennie P., Millbrae, CA, UNITED STATES

Pan, James, Belmont, CA, UNITED STATES

Paoni, Nicholas F., Belmont, CA, UNITED STATES Roy, Margaret Ann, San Francisco, CA, UNITED STATES Stewart, Timothy A., San Francisco, CA, UNITED STATES

Tumas, Daniel, Orinda, CA, UNITED STATES

Williams, P. Mickey, Half Moon Bay, CA, UNITED STATES

Wood, Wlliam I., Hillsborough, CA, UNITED STATES

Genentech, Inc. (U.S. corporation)

PATENT ASSIGNEE(S):

KIND NUMBER DATE -----

PATENT INFORMATION: APPLICATION INFO.:

US 2002192659 A1 20021219 US 2001-902853 A1 20010710

RELATED APPLN. INFO.:

Continuation of Ser. No. US 2000-665350, filed on 18

Sep 2000, PENDING

NUMBER DATE 19980910

PRIORITY INFORMATION: WO 1998-US18824

> WO 1998-US19177 19980914 WO 1998-US19330 19980916

WO 1998-US19437 19980917

WO 1998-US25108 19981201

WO 1999-US20594 19990908

WO 1999-US20944 19990913 WO 1999-US21090 19990915

WO 1999-US21547 19990915

WO 1999-US23089 19991005

WO 1999-US28214 19991129

WO 1999-US28313 19991130

WO 1999-US28301 19991201

WO 1999-US28564 19991202 19991202

WO 1999-US28565 WO 1999-US30095 19991216

WO 1999-US30999 19991220

WO 1999-US30911 19991220

WO 2000-US219 20000105 20000211

WO 2000-US3565 WO 2000-US4414 20000222

WO 2000-US5004 20000224

WO 2000-US5841 20000302

WO 2000-US7377 20000320

```
20000330
                         WO 2000-US8439
                         WO 2000-US14042
                                             20000522
                         WO 2000-US15264
                                             20000602
                         WO 2000-US20710
                                             20000728
                         WO 2000-US23328
                                             20000824
                         US 1997-59115P
                                             19970917 (60)
                                             19970917 (60)
                         US 1997-59184P
                         US 1997-59122P
                                             19970917
                                                      (60)
                         US 1997-59117P
                                             19970917
                                                      (60)
                         US 1997-59113P
                                             19970917
                                                      (60)
                         US 1997-59121P
                                             19970917
                                                      (60)
                         US 1997-59119P
                                             19970917
                                                      (60)
                         US 1997-59263P
                                             19970918 (60)
                         US 1997-59266P
                                             19970918 (60)
                                             19971015
                                                      (60)
                         US 1997-62125P
                                             19971017
                                                      (60)
                         US 1997-62287P
                         US 1997-62285P
                                             19971017
                                                      (60)
                         US 1997-63486P
                                             19971021 (60)
                         US 1997-62816P
                                             19971024 (60)
                         US 1997-62814P
                                             19971024 (60)
                         US 1997-63127P
                                             19971024 (60)
                         US 1997-63120P
                                             19971024 (60)
                         US 1997-63121P
                                             19971024 (60)
                         US 1997-63045P
                                             19971024 (60)
                         US 1997-63128P
                                             19971024 (60)
                         US 1997-63329P
                                             19971027
                         US 1997-63327P
                                             19971027
                         US 1997-63549P
                                             19971028
                         US 1997-63541P
                                             19971028
                         US 1997-63550P
                                             19971028
                         US 1997-63542P
                                             19971028
                         US 1997-63544P
                                             19971028 (60)
                         US 1997-63564P
                                             19971028 (60)
                         US 1997-63734P
                                             19971029 (60)
                         US 1997-63738P
                                             19971029 (60)
                         US 1997-63704P
                                             19971029 (60)
                         US 1997-63435P
                                             19971029 (60)
                         US 1997-64215P
                                             19971029 (60)
                         US 1997-63735P
                                             19971029 (60)
                         US 1997-63732P
                                            19971029 (60)
                         US 1997-64103P
                                            19971031 (60)
                         US 1997-63870P
                                             19971031 (60)
                         US 1997-64248P
                                             19971103 (60)
                         US 1997-64809P
                                             19971107 (60)
                         US 1997-65186P
                                             19971112 (60)
                         US 1997-65846P
                                             19971117 (60)
                         US 1997-65693P
                                             19971118 (60)
                         US 1997-66120P
                                             19971121 (60)
                         US 1997-66364P
                                             19971121 (60)
                         US 1997-66772P
                                             19971124 (60)
                         US 1997-66466P
                                             19971124 (60)
                         US 1997-66770P
                                             19971124 (60)
                         US 1997-66511P
                                             19971124 (60)
                                             19971124 (60)
                         US 1997-66453P
                        Utility
                        APPLICATION
                        BRINKS HOFER GILSON & LIONE, P.O. BOX 10395, CHICAGO,
                         IL, 60610
                         38
                         1
                         124 Drawing Page(s)
                         21726
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
```

DOCUMENT TYPE:

FILE SEGMENT:

LINE COUNT:

NUMBER OF CLAIMS:

EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

LEGAL REPRESENTATIVE:

L9 ANSWER 9 OF 44 USPATFULL

TI Peptide scaffold encapsulation of tissue cells and uses thereof

AB The invention features peptide scaffolds that are useful in the repair and replacement of various tissues. The invention also provides methods for making these scaffolds and methods for using them.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2002:287608 USPATFULL

TITLE: Peptide scaffold encapsulation of tissue cells and uses

thereof

INVENTOR(S): Kisiday, John, Watertown, MA, UNITED STATES

Grodzinsky, Alan, Lexington, MA, UNITED STATES Zhang, Shuguang, Lexington, MA, UNITED STATES

NUMBER KIND DATE
----ON: US 2002160471 A1 20021031

PATENT INFORMATION: US 2002160471 A1 20021031 APPLICATION INFO.: US 2001-778200 A1 20010206 (9)

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: CLARK & ELBING LLP, 101 FEDERAL STREET, BOSTON, MA,

02110

NUMBER OF CLAIMS: 18 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 9 Drawing Page(s)

LINE COUNT: 1010

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L9 ANSWER 10 OF 44 USPATFULL

TI Secreted and transmembrane polypeptides and nucleic acids encoding the

The present invention is directed to novel polypeptides and to nucleic acid molecules encoding those polypeptides. Also provided herein are vectors and host cells comprising those nucleic acid sequences, chimeric polypeptide molecules comprising the polypeptides of the present invention fused to heterologous polypeptide sequences, antibodies which bind to the polypeptides of the present invention and to methods for producing the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2002:287511 USPATFULL

TITLE: Secreted and transmembrane polypeptides and nucleic

acids encoding the same

INVENTOR(S): Ashkenazi, Avi, San Mateo, CA, UNITED STATES

Botstein, David, Belmont, CA, UNITED STATES
Desnoyers, Luc, San Francisco, CA, UNITED STATES
Eaton, Dan L., San Rafael, CA, UNITED STATES

Ferrara, Napoleone, San Francisco, CA, UNITED STATES Filvaroff, Ellen, San Francisco, CA, UNITED STATES

Fong, Sherman, Alameda, CA, UNITED STATES Gao, Wei-Qiang, Palo Alto, CA, UNITED STATES

Gerber, Hanspeter, San Francisco, CA, UNITED STATES Gerritsen, Mary E., San Mateo, CA, UNITED STATES Goddard, Audrey, San Francisco, CA, UNITED STATES Godowski, Paul J., Burlingame, CA, UNITED STATES Grimaldi, J. Christopher, San Francisco, CA, UNITED

STATES

Gurney, Austin L., Belmont, CA, UNITED STATES Hillan, Kenneth J., San Francisco, CA, UNITED STATES Kljavin, Ivar J., Lafayette, CA, UNITED STATES Mather, Jennie P., Millbrae, CA, UNITED STATES

Pan, James, Belmont, CA, UNITED STATES

Paoni, Nicholas F., Belmont, CA, UNITED STATES Roy, Margaret Ann, San Francisco, CA, UNITED STATES Stewart, Timothy A., San Francisco, CA, UNITED STATES Tumas, Daniel, Orinda, CA, UNITED STATES Williams, P. Mickey, Half Moon Bay, CA, UNITED STATES Wood, William I., Hillsborough, CA, UNITED STATES Genentech, Inc. (U.S. corporation)

PATENT ASSIGNEE(S):

KIND DATE NUMBER -----US 2002160374 A1 20021031 US 2001-905291 A1 20010712 (9)

PATENT INFORMATION: APPLICATION INFO.: RELATED APPLN. INFO.:

Continuation of Ser. No. US 2000-665350, filed on 18

DATE

Sep 2000, PENDING

NUMBER

# PRIORITY INFORMATION:

	NUMBER	DATE	
WO	1998-US18824	19980910	
WO	1998-US19177	19980914	
WO	1998-US19330	19980916	
		19980917	
WO	1998-US19437		
WO	1998-US25108	19981201	
WO	1999-US20594	19990908	
WO	1999-US20944	19990913	
WO	1999-US21090	19990915	
WO	1999-US21547	19990915	
WO	1999-US23089	19991005	
WO	1999-US28214	19991129	
WO	1999-US28313	19991130	
WO	1999-US28301	19991201	
WO	1999-US28564	19991202	
WO	1999-US28565	19991202	
WO	1999-US30095	19991216	
WO	1999-US30999	19991220	
WO	1999-US30911	19991220	
WO	2000-US219	20000105	
WO	2000-US3565	20000211	
WO	2000-US4414	20000222	
WO	2000-US5004	20000224	
WO	2000-US5841	20000302	
WO	2000-US7377	20000320	
WO	2000-US8439	20000330	
WO	2000-US14042	20000522	
WO	2000-US15264	20000602	
WO	2000-US20710	20000728	
WO	2000-US23328	20000824	
US	1997-59115P	19970917	(60)
US	1997-59184P	19970917	(60)
US	1997-59122P	19970917	(60)
US	1997-59117P	19970917	(60)
US	1997-59113P	19970917	(60)
US	1997-59121P	19970917	(60)
US	1997-59119P	19970917	(60)
US	1997-59119P	19970917	(60)
US	1997-59266P	19970918	(60)
US	1997-62125P	19971015	(60)
US	1997-62287P	19971017	(60)
US	1997-62285P	19971017	(60)
US	1997-63486P	19971021	(60)
US	1997-62816P	19971024	(60)
US	1997-62814P	19971024	(60)
US	1997-63127P	19971024	(60)
US	1997-63120P	19971024	(60)
US	1997-63121P	19971024	(60)
US	1997-63045P	19971024	(60)
US	1997-63128P	19971024	(60)

```
US 1997-63329P
                   19971027 (60)
US 1997-63327P
                  19971027 (60)
US 1997-63549P
                  19971028 (60)
US 1997-63541P
                  19971028 (60)
US 1997-63550P
                  19971028 (60)
US 1997-63542P
                  19971028 (60)
US 1997-63544P
                  19971028 (60)
US 1997-63564P
                  19971028 (60)
US 1997-63734P
                  19971029 (60)
US 1997-63738P
                   19971029 (60)
US 1997-63704P
                   19971029 (60)
                   19971029 (60)
US 1997-63435P
                   19971029 (60)
US 1997-64215P
                   19971029 (60)
US 1997-63735P
                   19971029 (60)
US 1997-63732P
                   19971031 (60)
US 1997-64103P
                   19971031 (60)
US 1997-63870P
                   19971103 (60)
US 1997-64248P
                   19971107 (60)
US 1997-64809P
US 1997-65186P
                   19971112 (60)
                   19971117 (60)
US 1997-65846P
                   19971118 (60)
US 1997-65693P
                   19971121 (60)
US 1997-66120P
                   19971121 (60)
US 1997-66364P
                   19971124 (60)
US 1997-66772P
                   19971124 (60)
US 1997-66466P
                   19971124 (60)
US 1997-66770P
US 1997-66511P
                   19971124 (60)
                   19971124 (60)
US 1997-66453P
Utility
```

DOCUMENT TYPE:

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

BRINKS HOFER GILSON & LIONE, P.O. BOX 10395, CHICAGO,

IL, 60610

NUMBER OF CLAIMS:

EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

124 Drawing Page(s)

LINE COUNT:

21310

38

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

#### ANSWER 11 OF 44 USPATFULL L9

Secreted and transmembrane polypeptides and nucleic acids encoding the TI

The present invention is directed to novel polypeptides and to nucleic AΒ acid molecules encoding those polypeptides. Also provided herein are vectors and host cells comprising those nucleic acid sequences, chimeric polypeptide molecules comprising the polypeptides of the present invention fused to heterologous polypeptide sequences, antibodies which bind to the polypeptides of the present invention and to methods for producing the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2002:265833 USPATFULL

TITLE:

Secreted and transmembrane polypeptides and nucleic

acids encoding the same

INVENTOR(S):

Ashkenazi, Avi, San Mateo, CA, UNITED STATES Botstein, David, Belmont, CA, UNITED STATES Desnoyers, Luc, San Francisco, CA, UNITED STATES Eaton, Dan L., San Rafael, CA, UNITED STATES Ferrara, Napoleone, San Francisco, CA, UNITED STATES Filvaroff, Ellen, San Francisco, CA, UNITED STATES Fong, Sherman, Alameda, CA, UNITED STATES Gao, Wei-Qiang, Palo Alto, CA, UNITED STATES Gerber, Hanspeter, San Francisco, CA, UNITED STATES Gerritsen, Mary E., San Mateo, CA, UNITED STATES

Goddard, Audrey, San Francisco, CA, UNITED STATES Godowski, Paul J., Burlingame, CA, UNITED STATES Grimaldi, J. Christopher, San Francisco, CA, UNITED STATES

Gurney, Austin L., Belmont, CA, UNITED STATES
Hillan, Kenneth J., San Francisco, CA, UNITED STATES
Kljavin, Ivar J., Lafayette, CA, UNITED STATES
Mather, Jennie P., Millbrae, CA, UNITED STATES
Pan, James, Belmont, CA, UNITED STATES
Paoni, Nicholas F., Belmont, CA, UNITED STATES

Roy, Margaret Ann, San Francisco, CA, UNITED STATES Stewart, Timothy A., San Francisco, CA, UNITED STATES Tumas, Daniel, Orinda, CA, UNITED STATES

Williams, P. Mickey, Half Moon Bay, CA, UNITED STATES Wood, William I., Hillsborough, CA, UNITED STATES Genentech, Inc. (U.S. corporation)

PATENT ASSIGNEE(S):

NUMBER KIND DATE

PATENT INFORMATION: APPLICATION INFO.: RELATED APPLN. INFO.: US 2002146709 A1 20021010 US 2001-909088 A1 20010718 (9)

Continuation of Ser. No. US 2000-665350, filed on 18 Sep 2000, PENDING

NUMBER DATE

## PRIORITY INFORMATION:

	MONDER	DATE	
	1998-US18824		
WO	1998-US19177	19980914	
WO	1998-US19330	19980916	
WO	1998-US19437	19980917	
WO	1998-US25108	19981201	
WO	1999-US20594	19990908	
WO	1999-US20944	19990913	
WO	1999-US21090	19990915	
WO	1999-US21547	19990915	
WO	1999-US23089	19991005	
WO	1999-US28214	19991129	
WO	1999-US28313	19991130	
WO	1999-US28301	19991201	
WO	1999-US28564	19991202	
WO	1999-US28565	19991202	
WO	1999-US30095	19991216	
WO	1999-US30999	19991220	
WO	1999-US30911	19991220	
WO	2000-US219	20000105	
WO	2000-US3565	20000211	
WO	2000-US4414	20000222	
WO	2000-US5004	20000224	
WO	2000-US5841	20000302	
WO	2000-US7377	20000320	
WO	2000-US8439	20000330	
WO	2000-US14042	20000522	
WO	2000-US15264	20000602	
WO	2000-US20710	20000728	
WO	2000-US23328	20000824	
US	1997-59115P	19970917	(60)
US	1997-59184P	19970917	(60)
US	1997-59122P	19970917	(60)
US	1997-59117P	19970917	(60)
US	1997-59113P	19970917	(60)
US	1997-59121P	19970917	(60)
US	1997-59119P	19970917	(60)
US	1997-59263P	19970918	(60)
US	1997-59266P	19970918	(60)

```
19971015 (60)
US 1997-62125P
                   19971017 (60)
US 1997-62287P
US 1997-62285P
                   19971017 (60)
US 1997-63486P
                   19971021 (60)
US 1997-62816P
                   19971024 (60)
                   19971024 (60)
US 1997-62814P
US 1997-63127P
                   19971024 (60)
US 1997-63120P
                   19971024 (60)
US 1997-63121P
                   19971024 (60)
US 1997-63045P
                   19971024 (60)
                   19971024 (60)
US 1997-63128P
                   19971027 (60)
US 1997-63329P
                   19971027 (60)
US 1997-63327P
                   19971028 (60)
US 1997-63549P
US 1997-63541P
                   19971028 (60)
US 1997-63550P
                   19971028 (60)
                   19971028 (60)
US 1997-63542P
                   19971028 (60)
US 1997-63544P
                   19971028 (60)
US 1997-63564P
                   19971029 (60)
US 1997-63734P
US 1997-63738P
                   19971029 (60)
US 1997-63704P
                   19971029 (60)
US 1997-63435P
                   19971029 (60)
US 1997-64215P
                   19971029 (60)
                   19971029 (60)
US 1997-63735P
                   19971029 (60)
US 1997-63732P
                   19971031 (60)
US 1997-64103P
                   19971031 (60)
US 1997-63870P
                   19971103 (60)
US 1997-64248P
                   19971107 (60)
US 1997-64809P
                   19971112 (60)
US 1997-65186P
                   19971117 (60)
US 1997-65846P
US 1997-65693P
                   19971118 (60)
US 1997-66120P
                   19971121 (60)
US 1997-66364P
                   19971121 (60)
US 1997-66772P
                   19971124 (60)
US 1997-66466P
                   19971124 (60)
US 1997-66770P
                   19971124 (60)
US 1997-66511P
                   19971124 (60)
US 1997-66453P
                   19971124 (60)
Utility
APPLICATION
```

DOCUMENT TYPE: FILE SEGMENT:

LEGAL REPRESENTATIVE:

BRINKS HOFER GILSON & LIONE, P.O. BOX 10395, CHICAGO,

IL, 60610

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

38

NUMBER OF DRAWINGS:

124 Drawing Page(s)

LINE COUNT: 21668

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L9 ANSWER 12 OF 44 USPATFULL

ΤI Bone morphogenic protein (BMP)

polynucleotides, polypeptides, and antibodies

The present invention relates to novel human BMP polypeptides and AB isolated nucleic acids containing the coding regions of the genes encoding such polypeptides. Also provided are vectors, host cells, antibodies, and recombinant methods for producing human BMP polypeptides. The invention further relates to diagnostic and therapeutic methods useful for diagnosing and treating disorders related to these novel human BMP polypeptides.

CAS INDEXING IS AVAILABLE FOR THIS PATENT. 2002:259593 USPATFULL ACCESSION NUMBER: Bone morphogenic protein

TITLE:

(BMP) polynucleotides, polypeptides, and antibodies

Ni, Jian, Germantown, MD, UNITED STATES Ruben, Steven M., Olney, MD, UNITED STATES Shi, Yanggu, Gaithersburg, MD, UNITED STATES

PATENT ASSIGNEE(S): Human Genome Sciences, Inc., Rockville, MD, UNITED

STATES, 20850 (U.S. corporation)

RELATED APPLN. INFO.: Continuation of Ser. No. US 2000-685899, filed on 11

Oct 2000, PENDING Continuation-in-part of Ser. No. WO

2000-US9028, filed on 6 Apr 2000, UNKNOWN

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 22 EXEMPLARY CLAIM: 1 LINE COUNT: 10845

INVENTOR(S):

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L9 ANSWER 13 OF 44 USPATFULL

TI Secreted and transmembrane polypeptides and nucleic acids encoding the

The present invention is directed to novel polypeptides and to nucleic acid molecules encoding those polypeptides. Also provided herein are vectors and host cells comprising those nucleic acid sequences, chimeric polypeptide molecules comprising the polypeptides of the present invention fused to heterologous polypeptide sequences, antibodies which bind to the polypeptides of the present invention and to methods for producing the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2002:243054 USPATFULL

TITLE: Secreted and transmembrane polypeptides and nucleic

acids encoding the same

INVENTOR(S): Ashkenazi, Avi, San Mateo, CA, UNITED STATES

Botstein, David, Belmont, CA, UNITED STATES
Desnoyers, Luc, San Francisco, CA, UNITED STATES
Eaton, Dan L., San Rafael, CA, UNITED STATES

Ferrara, Napoleone, San Francisco, CA, UNITED STATES

Filvaroff, Ellen, San Francisco, CA, UNITED STATES Fong, Sherman, Alameda, CA, UNITED STATES Gao, Wei-Qiang, Palo Alto, CA, UNITED STATES

Gerber, Hanspeter, San Francisco, CA, UNITED STATES Gerritsen, Mary E., San Mateo, CA, UNITED STATES Goddard, Audrey, San Francisco, CA, UNITED STATES Godowski, Paul J., Burlingame, CA, UNITED STATES Grimaldi, J. Christopher, San Francisco, CA, UNITED

STATES

Gurney, Austin L., Belmont, CA, UNITED STATES
Hillan, Kenneth J., San Francisco, CA, UNITED STATES

Kljavin, Ivar J., Lafayette, CA, UNITED STATES Mather, Jennie P., Millbrae, CA, UNITED STATES

Pan, James, Belmont, CA, UNITED STATES

Paoni, Nicholas F., Belmont, CA, UNITED STATES Roy, Margaret Ann, San Francisco, CA, UNITED STATES Stewart, Timothy A., San Francisco, CA, UNITED STATES Tumas, Daniel, Orinda, CA, UNITED STATES Williams, P. Mickey, Half Moon Bay, CA, UNITED STATES Wood, William I., Hillsborough, CA, UNITED STATES Genentech, Inc. (U.S. corporation)

PATENT ASSIGNEE(S):

NUMBER KIND DATE \_\_\_\_\_\_ US 2002132240 A1 20020919 US 2001-909320 A1 20010718 (9)

PATENT INFORMATION: APPLICATION INFO.:

RELATED APPLN. INFO.:

Continuation of Ser. No. US 2000-665350, filed on 18

DATE

Sep 2000, PENDING

NUMBER

# PRIORITY INFORMATION:

	NOMBER	DAIL	
WO	1998-US18824	19980910	
WO	1998-US19177		
WO	1998-US19330	19980916	
WO	1998-US19437	19980917	
WO	1998-US25108	19981201	
WO	1999-US20594	19990908	
WO	1999-US20944	19990913	
WO	1999-US21090	19990915	
WO	1999-US21547	19990915	
WO	1999-US23089	19991005	
WO	1999-US28214	19991129	
WO	1999-US28313	19991130	
WO	1999-US28301	19991201	
WO	1999-US28564	19991202	
WO	1999-US28565	19991202	
WO	1999-US30095	19991216	
WO	1999-US30999	19991220	
WO	1999-US30911	19991220	
WO	2000-US219	20000105	
WO	2000-US3565	20000211	
WO	2000-US4414	20000222	
WO	2000-US5004	20000224	
WO	2000-US5841	20000302	
WO	2000-US7377	20000320	
WO	2000-US8439	20000330	
WO	2000-US14042	20000522	
WO	2000-US15264	20000602	
WO	2000-US20710	20000728	
WO	2000-US23328	20000824	
US	1997-59115P	19970917	(60)
US	1997-59184P	19970917	(60)
US	1997-59122P	19970917	(60)
US	1997-59117P	19970917	(60)
US	1997-59113P	19970917	(60)
US	1997-59121P	19970917	(60)
US	1997-59119P	19970917	(60)
US	1997-59263P	19970918	(60)
US	1997-59266P	19970918	(60)
US	1997-62125P	19971015	(60)
US	1997-62287P	19971017	(60)
US	1997-62285P	19971017	(60)
US	1997-63486P	19971021	(60)
US	1997-62816P	19971024	(60)
US	1997-62814P	19971024	(60)
US	1997-63127P	19971024	
US	1997-63120P	19971024	
US	1997-63121P	19971024	
00	IJJI UJILIE	10011024	(00)

```
US 1997-63045P
                   19971024 (60)
                   19971024 (60)
US 1997-63128P
US 1997-63329P
                   19971027 (60)
                   19971027 (60)
US 1997-63327P
US 1997-63549P
                   19971028 (60)
US 1997-63541P
                   19971028 (60)
US 1997-63550P
                   19971028 (60)
US 1997-63542P
                   19971028 (60)
US 1997-63544P
                   19971028 (60)
US 1997-63564P
                   19971028 (60)
US 1997-63734P
                   19971029 (60)
US 1997-63738P
                   19971029 (60)
                   19971029 (60)
US 1997-63704P
                   19971029 (60)
US 1997-63435P
US 1997-64215P
                   19971029 (60)
                   19971029 (60)
US 1997-63735P
                   19971029 (60)
US 1997-63732P
                   19971031 (60)
US 1997-64103P
                   19971031 (60)
US 1997-63870P
                   19971103 (60)
US 1997-64248P
                   19971107 (60)
US 1997-64809P
                   19971112 (60)
US 1997-65186P
                   19971117 (60)
US 1997-65846P
                   19971118 (60)
US 1997-65693P
                   19971121 (60)
US 1997-66120P
                   19971121 (60)
US 1997-66364P
                   19971124 (60)
US 1997-66772P
                   19971124 (60)
US 1997-66466P
                   19971124 (60)
US 1997-66770P
                   19971124 (60)
US 1997-66511P
                   19971124 (60)
US 1997-66453P
```

DOCUMENT TYPE:

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

BRINKS HOFER GILSON & LIONE, P.O. BOX 10395, CHICAGO,

IL, 60610

Utility

NUMBER OF CLAIMS:

EXEMPLARY CLAIM:

38

NUMBER OF DRAWINGS:

124 Drawing Page(s)

LINE COUNT:

21778

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 14 OF 44 USPATFULL

Use of adipose tissue-derived stromal cells for chondrocyte TТ

differentiation and cartilage repair

AB Methods and compositions for directing adipose-derived stromal cells cultivated in vitro to differentiate into cells of the chondrocyte lineage are disclosed. The invention further provides a variety of chondroinductive agents which can be used singly or in combination with other nutrient components to induce chondrogenesis in adipose-derived stromal cells either in cultivating monolayers or in a biocompatible lattice or matrix in a three-dimensional configuration. Use of the differentiated chondrocytes for the therapeutic treatment of a number of human conditions and diseases including repair of cartilage in vivo is disclosed.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

2002:214259 USPATFULL ACCESSION NUMBER:

Use of adipose tissue-derived stromal cells for TITLE:

chondrocyte differentiation and cartilage repair

Halvorsen, Yuan-Di C., Holly Springs, NC, UNITED STATES INVENTOR(S):

Wilkison, William O., Bahama, NC, UNITED STATES

Gimble, Jeffrey Martin, Chapel Hill, NC, UNITED STATES

NUMBER KIND DATE -----

PATENT INFORMATION: US 2002115647 A1 20020822 APPLICATION INFO.: US 2002-125106 A1 20020418 (10) APPLICATION INFO.:

Continuation of Ser. No. US 2000-573989, filed on 17 RELATED APPLN. INFO.:

May 2000, PENDING

NUMBER DATE

\_\_\_\_\_ US 1999-149850P 19990819 (60) PRIORITY INFORMATION:

DOCUMENT TYPE: Utility

APPLICATION FILE SEGMENT:

KING & SPALDING, 191 PEACHTREE STREET, N.E., ATLANTA, LEGAL REPRESENTATIVE:

GA, 30303-1763

NUMBER OF CLAIMS: 18 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 5 Drawing Page(s)

831 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 15 OF 44 USPATFULL

Use of adipose tissue-derived stromal cells for chondrocyte ΤI

differentiation and cartilage repair

AB Methods and compositions for directing adipose-derived stromal cells cultivated in vitro to differentiate into cells of the chondrocyte lineage are disclosed. The invention further provides a variety of chondroinductive agents which can be used singly or in combination with other nutrient components to induce chondrogenesis in adipose-derived stromal cells either in cultivating monolayers or in a biocompatible lattice or matrix in a three-dimensional configuration. Use of the differentiated chondrocytes for the therapeutic treatment of a number of human conditions and diseases including repair of cartilage in vivo is disclosed.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2002:194742 USPATFULL

Use of adipose tissue-derived stromal cells for TITLE:

chondrocyte differentiation and cartilage repair

Halvorsen, Yuan-Di C., Holly Springs, NC, United States INVENTOR(S):

Wilkison, William O., Bahama, NC, United States

Gimble, Jeffrey Martin, Chapel Hill, NC, United States

Artecel Science, Inc., Durham, NC, United States (U.S. PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 6429013 B1 20020806 US 2000-573989 20000517 APPLICATION INFO.: 20000517 (9)

> NUMBER DATE \_\_\_\_\_

PRIORITY INFORMATION: US 1999-149850P 19990819 (60)

DOCUMENT TYPE: Utility GRANTED FILE SEGMENT: Guzo, David PRIMARY EXAMINER:

ASSISTANT EXAMINER: Davis, Katharine F

King & Spalding, Knowles, Sherry M., Bennett-Paris, LEGAL REPRESENTATIVE:

Joseph

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

5 Drawing Figure(s); 5 Drawing Page(s) NUMBER OF DRAWINGS:

LINE COUNT: 995

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Welcome to STN International! Enter x:x

```
LOGINID:ssspta1653hxp
PASSWORD:
TERMINAL (ENTER 1, 2, 3, OR ?):2
                      Welcome to STN International
                  Web Page URLs for STN Seminar Schedule - N. America
 NEWS
                  "Ask CAS" for self-help around the clock
 NEWS
          Apr 08
                  BEILSTEIN: Reload and Implementation of a New Subject Area
 NEWS
          Apr 09
                  ZDB will be removed from STN
 NEWS
          Apr 09
                  US Patent Applications available in IFICDB, IFIPAT, and IFIUDB
          Apr 19
 NEWS
                  Records from IP.com available in CAPLUS, HCAPLUS, and ZCAPLUS
 NEWS
          Apr 22
                  BIOSIS Gene Names now available in TOXCENTER
 NEWS
          Apr 22
                  Federal Research in Progress (FEDRIP) now available
 NEWS
          Apr 22
                  New e-mail delivery for search results now available
          Jun 03
 NEWS
         Jun 10
                  MEDLINE Reload
 NEWS 10
          Jun 10
                  PCTFULL has been reloaded
 NEWS 11
                  FOREGE no longer contains STANDARDS file segment
          Jul 02
 NEWS 12
                  USAN to be reloaded July 28, 2002;
 NEWS 13
         Jul 22
                  saved answer sets no longer valid
                  Enhanced polymer searching in REGISTRY
          Jul 29
 NEWS 14
          Jul 30
                  NETFIRST to be removed from STN
 NEWS 15
          Aug 08
                  CANCERLIT reload
 NEWS 16
                  PHARMAMarketLetter(PHARMAML) - new on STN
 NEWS 17
          Aug 08
                  NTIS has been reloaded and enhanced
          Aug 08
 NEWS 18
                  Aquatic Toxicity Information Retrieval (AQUIRE)
 NEWS 19
          Aug 19
                  now available on STN
                  IFIPAT, IFICDB, and IFIUDB have been reloaded
 NEWS 20
          Aug 19
                  The MEDLINE file segment of TOXCENTER has been reloaded
 NEWS 21
          Aug 19
 NEWS 22
          Aug 26
                  Sequence searching in REGISTRY enhanced
                  JAPIO has been reloaded and enhanced
 NEWS 23
          Sep 03
                  Experimental properties added to the REGISTRY file
          Sep 16
 NEWS 24
                  CA Section Thesaurus available in CAPLUS and CA
 NEWS 25
          Sep 16
                  CASREACT Enriched with Reactions from 1907 to 1985
 NEWS 26
         Oct 01
 NEWS 27
         Oct 21
                  EVENTLINE has been reloaded
                  BEILSTEIN adds new search fields
 NEWS 28 Oct 24
                  Nutraceuticals International (NUTRACEUT) now available on STN
 NEWS 29 Oct 24
                  MEDLINE SDI run of October 8, 2002
 NEWS 30 Oct 25
 NEWS 31 Nov 18
                  DKILIT has been renamed APOLLIT
                  More calculated properties added to REGISTRY
 NEWS 32 Nov 25
 NEWS 33
         Dec 02
                  TIBKAT will be removed from STN
         Dec 04
                  CSA files on STN
 NEWS 34
                  PCTFULL now covers WP/PCT Applications from 1978 to date
 NEWS 35
         Dec 17
                  TOXCENTER enhanced with additional content
         Dec 17
 NEWS 36
                  Adis Clinical Trials Insight now available on STN
          Dec 17
 NEWS 37
 NEWS 38
          Dec 30
                  ISMEC no longer available
                  Indexing added to some pre-1967 records in CA/CAPLUS
 NEWS 39
          Jan 13
               January 6 CURRENT WINDOWS VERSION IS V6.01a,
 NEWS EXPRESS
               CURRENT MACINTOSH VERSION IS V6.0b(ENG) AND V6.0Jb(JP),
               AND CURRENT DISCOVER FILE IS DATED 01 OCTOBER 2002
               STN Operating Hours Plus Help Desk Availability
 NEWS HOURS
               General Internet Information
 NEWS INTER
               Welcome Banner and News Items
 NEWS LOGIN
```

NEWS PHONE Direct Dial and Telecommunication Network Access to STN NEWS WWW CAS World Wide Web Site (general information)

Enter NEWS followed by the item number or name to see news on that specific topic.

All use of STN is subject to the provisions of the STN Customer agreement. Please note that this agreement limits use to scientific research. Use for software development or design or implementation of commercial gateways or other similar uses is prohibited and may result in loss of user privileges and other penalties.

FILE 'HOME' ENTERED AT 10:30:42 ON 15 JAN 2003

=> file medline, dgene, embase, scisearch, uspatful, wpids, jicst, fsta
COST IN U.S. DOLLARS

SINCE FILE
ENTRY
SESSION
FULL ESTIMATED COST

0.42
0.42

FILE 'MEDLINE' ENTERED AT 10:31:56 ON 15 JAN 2003

FILE 'DGENE' ENTERED AT 10:31:56 ON 15 JAN 2003 COPYRIGHT (C) 2003 THOMSON DERWENT

FILE 'EMBASE' ENTERED AT 10:31:56 ON 15 JAN 2003 COPYRIGHT (C) 2003 Elsevier Science B.V. All rights reserved.

FILE 'SCISEARCH' ENTERED AT 10:31:56 ON 15 JAN 2003 COPYRIGHT (C) 2003 Institute for Scientific Information (ISI) (R)

FILE 'USPATFULL' ENTERED AT 10:31:56 ON 15 JAN 2003 CA INDEXING COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'WPIDS' ENTERED AT 10:31:56 ON 15 JAN 2003 COPYRIGHT (C) 2003 THOMSON DERWENT

FILE 'JICST-EPLUS' ENTERED AT 10:31:56 ON 15 JAN 2003 COPYRIGHT (C) 2003 Japan Science and Technology Corporation (JST)

FILE 'FSTA' ENTERED AT 10:31:56 ON 15 JAN 2003 COPYRIGHT (C) 2003 International Food Information Service

=> s articular cartilage

L2 26968 ARTICULAR CARTILAGE

=> s l1 and l2 L5 76 L1 AND L2

=> s 15 and 13

L6 1 L5 AND L3

L6 ANSWER 1 OF 1 USPATFULL

TI Device for regeneration of articular cartilage and

other tissue

An implantable device for facilitating the healing of voids in bone, cartilage and soft tissue is disclosed. A preferred embodiment includes a cartilage region comprising a polyelectrolytic complex joined with a subchondral bone region. The cartilage region, of this embodiment, enhances the environment for chondrocytes to grow articular cartilage; while the subchondral bone region enhances the environment for cells which migrate into that region's macrostructure and which differentiate into osteoblasts. A hydrophobic barrier exists between said regions, of this embodiment. In one embodiment, the polyelectrolytic complex transforms to hydrogel, following the implant procedure.

ACCESSION NUMBER:

2002:55324 USPATFULL

TITLE:

Device for regeneration of articular

cartilage and other tissue

INVENTOR (S):

Brekke, John H., Duluth, MN, UNITED STATES Goldman, Scott M., Paoli, PA, UNITED STATES

NUMBER KIND DATE
----US 2002032488 A1 20020314
US 2001-909027 A1 20010719 (9)

PATENT INFORMATION: APPLICATION INFO.: RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 1998-206604, filed on 7 Dec 1998, GRANTED, Pat. No. US 6264701 Division of Ser. No. US 1994-242557, filed on 13 May 1994, GRANTED,

Pat. No. US 5981825

DOCUMENT TYPE: FILE SEGMENT:

Utility APPLICATION

LEGAL REPRESENTATIVE:

Alan D. Kamrath, Kensey Nash Corporation, 55 E. Uwchlan

Avenue, Exton, PA, 19341

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 56

LINE COUNT:

1349

## => d his

L1

L3

(FILE 'HOME' ENTERED AT 10:30:42 ON 15 JAN 2003)

FILE 'MEDLINE, DGENE, EMBASE, SCISEARCH, USPATFULL, WPIDS, JICST-EPLUS, FSTA' ENTERED AT 10:31:56 ON 15 JAN 2003

2316 S BONE MORPHOGENIC PROTEIN

L2 26968 S ARTICULAR CARTILAGE

125 S OSTEOCHONDRAL GRAFT

L4 0 S L1 () L2 () L3

L5 76 S L1 AND L2

L6 1 S L5 AND L3

=> s 13 and 11

L7 1 L3 AND L1

## => d 17 ti abs ibib tot

- L7 ANSWER 1 OF 1 USPATFULL
- TI Device for regeneration of articular cartilage and other tissue
- AB An implantable device for facilitating the healing of voids in bone, cartilage and soft tissue is disclosed. A preferred embodiment includes a cartilage region comprising a polyelectrolytic complex joined with a

subchondral bone region. The cartilage region, of this embodiment, enhances the environment for chondrocytes to grow articular cartilage; while the subchondral bone region enhances the environment for cells which migrate into that region's macrostructure and which differentiate into osteoblasts. A hydrophobic barrier exists between said regions, of this embodiment. In one embodiment, the polyelectrolytic complex transforms to hydrogel, following the implant procedure.

2002:55324 USPATFULL ACCESSION NUMBER:

Device for regeneration of articular cartilage and TITLE:

other tissue

Brekke, John H., Duluth, MN, UNITED STATES INVENTOR(S):

Goldman, Scott M., Paoli, PA, UNITED STATES

KIND DATE NUMBER \_\_\_\_\_\_

US 2002032488 A1 20020314 US 2001-909027 A1 20010719 (9) PATENT INFORMATION: APPLICATION INFO.:

Continuation-in-part of Ser. No. US 1998-206604, filed RELATED APPLN. INFO.:

on 7 Dec 1998, GRANTED, Pat. No. US 6264701 Division of Ser. No. US 1994-242557, filed on 13 May 1994, GRANTED,

Pat. No. US 5981825

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

Alan D. Kamrath, Kensey Nash Corporation, 55 E. Uwchlan LEGAL REPRESENTATIVE:

Avenue, Exton, PA, 19341

56 NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1349 LINE COUNT:

=> d his

(FILE 'HOME' ENTERED AT 10:30:42 ON 15 JAN 2003)

FILE 'MEDLINE, DGENE, EMBASE, SCISEARCH, USPATFULL, WPIDS, JICST-EPLUS, FSTA' ENTERED AT 10:31:56 ON 15 JAN 2003

2316 S BONE MORPHOGENIC PROTEIN

L1 L2 26968 S ARTICULAR CARTILAGE

125 S OSTEOCHONDRAL GRAFT L3

0 S L1 () L2 () L3 L4

76 S L1 AND L2 L5

1 S L5 AND L3 L6

L7 1 S L3 AND L1

=> d 15 ti abs ibib 1-10

ANSWER 1 OF 76 MEDLINE

Long-term effect of nitric oxide synthase inhibitor on repair of ΤI articular cartilage defects repairing.

OBJECTIVE: To discuss the long-term effect of inducible nitric oxide AB synthase inhibitor S-methylisothiourea (SMT) on repair of articular cartilage defects. METHODS: Twenty-four adult New Zealand White rabbits with full-thickness defects of cartilage created in the trochlear groove of their bilateral femurs were divided into three groups randomly, 8 in each group: (1) control group in which nothing was filled into the defects; (2) BMP group in which the defects were filled with collagen fibrin gel impregnated with recombinant human bone morphogenic protein (rhBMP); and (3) SMT group in which the defects were filled with collagen fibrin gel impregnated with rhBMP and hypodermic injection of SMT (5 mg .(-1) 12 h(-1)) was given. The animals were killed one year later. The gross appearance of the defects was assessed. The amount of released NO and the activity of NOS were examined by chemical colorimetry. The distribution of collagen was

examined by immunohistochemistry. The proteoglycan synthesis and cell activity was assessed by incorporation of radiolabelled sodium sulphate Na(2)(35)SO(4) and bromodeoxyuridine. RESULTS: One year after the defects in SMT group showed greater improvement in margin integration, cellular morphology, and architecture within defect than those in BMP group and control group (P < 0.01). Immunohistochemistry showed that there was less type-I collagen and more type-II collagen in SMT group than in the other two groups. Radiolabelled sodium sulphate (Na(2)(35)SO(4)) incorporation test showed that the proteoglycan synthesis in defects was higher in SMT group than in the other two groups (P < 0.01). BrdU incorporation test showed cells in repaired tissue with remarkable proliferous activity. CONCLUSION: iNOS inhibitor SMT significantly improves the quality of repair of defected cartilage and delays its degradation.

ACCESSION NUMBER:

2002215446

MEDLINE

DOCUMENT NUMBER:

21951165 PubMed ID: 11953121

TITLE:

Long-term effect of nitric oxide synthase inhibitor on

repair of articular cartilage defects

repairing.

AUTHOR:

Sun Wei; Wang Jixing; Jin Dadi; Liu Xiaoxia

CORPORATE SOURCE:

Department of Orthopaedics Surgery, Nanfang Hospital

Affiliated to First Militery Medical University, Guangzhou

510515, China.

SOURCE:

CHUNG-HUA I HSUEH TSA CHIH [CHINESE MEDICAL JOURNAL], (2002

Jan 10) 82 (1) 23-6.

Journal code: 7511141. ISSN: 0376-2491.

PUB. COUNTRY:

China

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

Chinese

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

200205

ENTRY DATE:

Entered STN: 20020416

Last Updated on STN: 20020515 Entered Medline: 20020514

L5 ANSWER 2 OF 76 MEDLINE

TI Enhanced matrix synthesis and in vitro formation of cartilage-like tissue by genetically modified chondrocytes expressing BMP-7.

Bone morphogenic protein-7 (BMP-7) supports ectopic cartilage and bone formation, is expressed in normal articular cartilage, and increases matrix synthesis in chondrocytes. Based on this knowledge, we hypothesized that an adenovirus (Ad) vector encoding human BMP-7 could be used to modify chondrocytes qenetically to improve their capacity for cartilage repair. An adenovirus vector encoding BMP-7 (AdBMP-7) was constructed and its bioactivity confirmed by ectopic bone formation assay. AdBMP-7 modification of bovine chondrocytes induced expression of BMP-7 mRNA and bioactive protein, resulting in an increase in incorporation of 35SO4- into proteoglycan, 3H-proline uptake into protein, and the expression of the cartilage-specific matrix genes, aggrecan and type II collagen. An in vitro model of chondrocyte transplantation was used to demonstrate the feasibility of using genetically modified chondrocytes to enhance formation of cartilage-like tissue. When transplanted onto cartilage explants and maintained in vitro for 3 weeks, chondrocytes modified with AdBMP-7 formed 1.9-fold thicker tissue than chondrocytes modified with a control vector (P < 0.001). This tissue was positive for type II collagen and proteoglycan but negative for type X collagen and demonstrated a cartilage-like morphology. These observations suggest that Ad-mediated transfer of BMP-7 gene to chondrocytes enhances the chondrocyte-specific matrix synthesis and their capacity to form cartilage-like tissue, thus representing a strategy that may improve cell-based cartilage repair.

ACCESSION NUMBER:

2001514436 MEDLINE

DOCUMENT NUMBER:

21446023 PubMed ID: 11562118

TITLE:

Enhanced matrix synthesis and in vitro formation of cartilage-like tissue by genetically modified chondrocytes

expressing BMP-7.

AUTHOR: Hidaka C; Quitoriano M; Warren R F; Crystal R G

CORPORATE SOURCE: Division of Pulmonary and Critical Care Medicine, Weill

Medical College of Cornell University, New York, NY, USA...

geneticmedicine@mail.med.cornell.edu

CONTRACT NUMBER: T32-AR07281 (NIAMS)

SOURCE: JOURNAL OF ORTHOPAEDIC RESEARCH, (2001 Sep) 19 (5) 751-8.

Journal code: 8404726. ISSN: 0736-0266.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200110

ENTRY DATE: Entered STN: 20010920

Last Updated on STN: 20011008 Entered Medline: 20011004

L5 ANSWER 3 OF 76 MEDLINE

TI Cartilage and bone regeneration using gene-enhanced tissue engineering.

Joint cartilage injury remains a major problem in orthopaedics with more than 500,000 cartilage repair procedures performed yearly in the United States at a cost of hundreds of millions of dollars. No consistently reliable means to regenerate joint cartilage currently exists. The technologies of gene therapy and tissue engineering were combined using a retroviral vector to stably introduce the human bone morphogenic protein-7 complementary deoxyribonucleic acid into periosteal-derived rabbit mesenchymal stem cells. Bone morphogenic protein-7 secreting gene modified cells subsequently were expanded in monolayer culture, seeded onto polyglycolic acid grafts, implanted into a rabbit knee osteochondral defect model, and evaluated for bone and cartilage repair after 4, 8, and 12 weeks. The grafts containing bone morphogenic protein-7

gene modified cells consistently showed complete or near complete bone and articular cartilage regeneration at 8 and 12 weeks

whereas the grafts from the control groups had poor repair as judged by macroscopic, histologic, and immunohistologic criteria. This is the first report of articular cartilage regeneration using a

combined gene therapy and tissue engineering approach.

ACCESSION NUMBER: 2000488818 MEDLINE

DOCUMENT NUMBER: 20492911 PubMed ID: 11039767

TITLE: Cartilage and bone regeneration using gene-enhanced tissue

engineering.

AUTHOR: Mason J M; Breitbart A S; Barcia M; Porti D; Pergolizzi R

G; Grande D A

CORPORATE SOURCE: Department of Research, North Shore University Hospital-New

York University School of Medicine, Manhasset 11030, USA.

SOURCE: CLINICAL ORTHOPAEDICS AND RELATED RESEARCH, (2000 Oct) (379

Suppl) S171-8.

Journal code: 0075674. ISSN: 0009-921X.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals

ENTRY MONTH: 200011

ENTRY DATE: Entered STN: 20010322

Last Updated on STN: 20010322 Entered Medline: 20001103

L5 ANSWER 4 OF 76 DGENE (C) 2003 THOMSON DERWENT

Isolated DNA encoding human SDF-5 protein - useful for controlling growth, differentiation etc. of cells, particularly of chondrocytes for treatment of arthritis etc., also pancreatic cells

AN AAW49082 Protein DGENE

AB The sequence is that of human SDF-5, a member of the Frazzled protein

family. Cells transformed with a vector containing the sequence are used to regulate genes, particularly pancreatic genes, or in combination with bone morphogenic protein 2 (BMP2), to

increase differentiation of progenitor cells into chondrocytes. The protein may be used to treat osteoarthritis, rheumatoid arthritis, or articular cartilage defects, also to increase/inhibit

articular cartilage defects, also to increase/inhibit cell formation, growth, differentiation, proliferation and/or maintenance in many other organs or tissues, e.g. for prevention or treatment of pancreatic cancer, diabetes (by inducing de novo formation of islet cells), other tissue defects, also to improve healing of wounds and to increase survival of nervous system cells, e.g. in cases of transplants The coding sequence can be used in gene therapy, and its fragments to detect related mRNA, while the protein is also used to generate antibodies, useful for affinity purification and as immunoassay reagents. Many other potential uses/activities for the gene and its encoded are contemplated but not exemplified, e.g. as cytokines, immuno-suppressants or immunostimulants, regulators of haematopoiesis, as fertility-control agents, haemostatic or thrombolytic agents, anti-inflammatory agents, antimicrobials, modulators of biorhythms and many more.

ACCESSION NUMBER: AAW49082 Protein DGENE

TITLE: Isolated DNA encoding human SDF-5 protein - useful for

controlling growth, differentiation etc. of cells,

particularly of chondrocytes for treatment of arthritis etc.,

69p

also pancreatic cells

INVENTOR: Lavallie E R; Racie L A
PATENT ASSIGNEE: (GEMY)GENETICS INST INC.
PATENT INFO: WO 9835043 A1 19980813

APPLICATION INFO: WO 1997-US18369 19971015 PRIORITY INFO: US 1997-848439 19970508

US 1997-796153 19970206

DOCUMENT TYPE: Patent LANGUAGE: English

OTHER SOURCE: 1998-447240 [38]

L5 ANSWER 5 OF 76 DGENE (C) 2003 THOMSON DERWENT

TI Isolated DNA encoding human SDF-5 protein - useful for controlling growth, differentiation etc. of cells, particularly of chondrocytes for treatment of arthritis etc., also pancreatic cells

AN AAV32930 DNA DGENE

The sequence is that encoding human SDF-5, a member of the Frazzled AB protein family. Cells transformed with a vector containing the sequence are used to regulate genes, particularly pancreatic genes, or in combination with bone morphogenic protein 2 (BMP2), to increase differentiation of progenitor cells into chondrocytes. The protein may be used to treat osteoarthritis, rheumatoid arthritis, or articular cartilage defects, also to increase/inhibit cell formation, growth, differentiation, proliferation and/or maintenance in many other organs or tissues, e.g. for prevention or treatment of pancreatic cancer, diabetes (by inducing de novo formation of islet cells), other tissue defects, also to improve healing of wounds and to increase survival of nervous system cells, e.g. in cases of transplants The coding sequence can be used in gene therapy, and its fragments to detect related mRNA, while the protein is also used to generate antibodies, useful for affinity purification and as immunoassay reagents. Many other potential uses/activities for the gene and its encoded are contemplated but not exemplified, e.g. as cytokines,

immuno-suppressants or immunostimulants, regulators of haematopoiesis, as fertility-control agents, haemostatic or thrombolytic agents, anti-inflammatory agents, antimicrobials, modulators of biorhythms and many more.

ACCESSION NUMBER: AAV32930 DNA DGENE

TITLE: Isolated DNA encoding human SDF-5 protein - useful for controlling growth, differentiation etc. of cells,

particularly of chondrocytes for treatment of arthritis etc.,

also pancreatic cells

INVENTOR: Lavallie E R; Racie L A
PATENT ASSIGNEE: (GEMY)GENETICS INST INC.
PATENT INFO: WO 9835043 A1 19980813

APPLICATION INFO: WO 1997-US18369 19971015 PRIORITY INFO: US 1997-848439 19970508

US 1997-796153 19970206

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 1998-447240 [38]

L5 ANSWER 6 OF 76 EMBASE COPYRIGHT 2003 ELSEVIER SCI. B.V.

TI Enhanced matrix synthesis and in vitro formation of cartilage-like tissue by genetically modified chondrocytes expressing BMP-7.

AB Bone morphogenic protein-7 (BMP-7) supports

ectopic cartilage and bone formation, is expressed in normal articular cartilage, and increases matrix synthesis in chondrocytes. Based on this knowledge, we hypothesized that an adenovirus (Ad) vector encoding human BMP-7 could be used to modify chondrocytes genetically to improve their capacity for cartilage repair. An adenovirus vector encoding BMP-7 (AdBMP-7) was constructed and its bioactivity confirmed by ectopic bone formation assay. AdBMP-7 modification of bovine chondrocytes induced expression of BMP-7 mRNA and bioactive protein, resulting in an increase in incorporation of (35)SO(-)(4) into proteoglycan, (3) H-proline uptake into protein, and the expression of the cartilage-specific matrix genes, aggrecan and type II collagen. An in vitro model of chondrocyte transplantation was used to demonstrate the feasibility of using genetically modified chondrocytes to enhance formation of cartilage-like tissue. When transplanted onto cartilage explants and maintained in vitro for 3 weeks, chondrocytes modified with AdBMP-7 formed 1.9-fold thicker tissue than chondrocytes modified with a control vector (P < 0.001). This tissue was positive for type II collagen and proteoglycan but negative for type X collagen and demonstrated a cartilage-like morphology. These observations suggest that Ad-mediated transfer of BMP-7 gene to chondrocytes enhances the chondrocyte-specific matrix synthesis and their capacity to form cartilage-like tissue, thus representing a strategy that may improve cell-based cartilage repair. .COPYRGT. 2001. Orthopaedic Research Society. Published by Elsevier Science Ltd. All rights reserved.

ACCESSION NUMBER: 2001305182 EMBASE

ACCESSION NUMBER: 2001303102 EMBASE

TITLE: Enhanced matrix synthesis and in vitro formation of

cartilage-like tissue by genetically modified chondrocytes

expressing BMP-7.

AUTHOR: Hidaka C.; Quitoriano M.; Warren R.F.; Crystal R.G.

CORPORATE SOURCE: C. Hidaka, Institute of Genetic Medicine, Weill Medical

Coll. of Cornell Univ., New York, NY 10021, United States.

geneticmedicine@mail.med.cornell.edu

SOURCE: Journal of Orthopaedic Research, (2001) 19/5 (751-758).

Refs: 40

ISSN: 0736-0266 CODEN: JOREDR

PUBLISHER IDENT.: S 0736-0266(01)00019-5

COUNTRY:

DOCUMENT TYPE:

FILE SEGMENT:

United Kingdom
Journal; Article
004 Microbiology
022 Human Genetics

029 Clinical Biochemistry 033 Orthopedic Surgery

LANGUAGE: English SUMMARY LANGUAGE: English

L5 ANSWER 7 OF 76 EMBASE COPYRIGHT 2003 ELSEVIER SCI. B.V.

TI Cartilage and bone regeneration using gene-enhanced tissue engineering.

AB Joint cartilage injury remains a major problem in orthopaedics with more than 500,000 cartilage repair procedures performed yearly in the United

69p

States at a cost of hundreds of millions of dollars. No consistently reliable means to reqenerate joint cartilage currently exists. The technologies of gene therapy and tissue engineering were combined using a retroviral vector to stably introduce the human bone

morphogenic protein-7 complementary deoxyribonucleic

acid into periosteal-derived rabbit mesenchymal stem cells. Bone

morphogenic protein-7 secreting gene modified cells

subsequently were expanded in monolayer culture, seeded onto polyglycolic acid grafts, implanted into a rabbit knee osteochondral defect model, and evaluated for bone and cartilage repair after 4, 8, and 12 weeks. The grafts containing bone morphogenic protein-7

gene modified cells consistently showed complete or near complete bone and articular cartilage regeneration at 8 and 12 weeks

whereas the grafts from the control groups had poor repair as judged by macroscopic, histologic, and immunohistologic criteria. This is the first report of articular cartilage regeneration using a

combined gene therapy and tissue engineering approach.

ACCESSION NUMBER:

2000362559 EMBASE

TITLE:

Cartilage and bone regeneration using gene-enhanced tissue

engineering.

AUTHOR:

Mason J.M.; Breitbart A.S.; Barcia M.; Porti D.; Pergolizzi

R.G.; Grande D.A.

CORPORATE SOURCE:

Dr. J.M. Mason, Gene Therapy Vector Laboratory, Department of Research, North Shore University Hospital, 350 Community

Drive, Manhasset, NY 11030, United States

SOURCE:

Clinical Orthopaedics and Related Research, (2000) -/379

SUPPL. (S171-S178).

Refs: 27

ISSN: 0009-921X CODEN: CORTBR

COUNTRY:

United States

DOCUMENT TYPE:

Journal; Conference Article

FILE SEGMENT:

Human Genetics 022 Orthopedic Surgery 033

036 Health Policy, Economics and Management

LANGUAGE: English SUMMARY LANGUAGE: English

ANSWER 8 OF 76 SCISEARCH COPYRIGHT 2003 ISI (R) L5

Cartilage and bone regeneration using gene-enhanced tissue engineering ΤI AΒ Joint cartilage injury remains a major problem in orthopaedics with

more than 500,000 cartilage repair procedures performed yearly in the United States at a cost of hundreds of millions of dollars. No consistently reliable means to regenerate joint cartilage currently exists. The technologies of gene therapy and tissue engineering were combined using a retroviral vector to stably introduce the human hone morphogenic protein-7 complementary deoxyribonucleic acid into periosteal-derived rabbit mesenchymal stem cells. Bone

morphogenic protein-7 secreting gene modified cells

subsequently were expanded in monolayer culture, seeded onto polyglycolic acid grafts, implanted into a rabbit knee osteochondral defect model, and evaluated for bone and cartilage repair after 4, 8, and 12 weeks, The grafts containing bone morphogenic protein-7

gene modified cells consistently showed complete or near complete bone and articular cartilage regeneration at 8 and 12 weeks

whereas the grafts from the control groups had poor repair as judged by macroscopic, histologic, and immunohistologic criteria, This is the first report of articular cartilage regeneration using a combined gene therapy and tissue engineering approach.

2000:777821 SCISEARCH ACCESSION NUMBER:

THE GENUINE ARTICLE: 362NP

TITLE: Cartilage and bone regeneration using gene-enhanced tissue

engineering

Mason J M (Reprint); Breitbart A S; Barcia M; Porti D; AUTHOR:

Pergolizzi R G; Grande D A

NYU, GENE THERAPY VECTOR LAB, DEPT RES, N SHORE UNIV HOSP, CORPORATE SOURCE:

SCH MED, 350 COMMUNITY DR, MANHASSET, NY 11030 (Reprint); NYU, DIV PLAST & RECONSTRUCT SURG, SCH MED, N SHORE UNIV HOSP, MANHASSET, NY 11030; NYU, DIV ORTHOPED SURG, SCH MED, N SHORE UNIV HOSP, DEPT SURG, MANHASSET, NY 11030

COUNTRY OF AUTHOR: USA

SOURCE:

CLINICAL ORTHOPAEDICS AND RELATED RESEARCH, (OCT 2000) No.

379, Supp. [S], pp. S171-S178.

Publisher: LIPPINCOTT WILLIAMS & WILKINS, 530 WALNUT ST,

PHILADELPHIA, PA 19106-3621.

ISSN: 0009-921X. Article; Journal

DOCUMENT TYPE: FILE SEGMENT:

LIFE; CLIN

LANGUAGE:

REFERENCE COUNT:

English

27

\*ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS\*

ANSWER 9 OF 76 USPATFULL L5

Secreted and transmembrane polypeptides and nucleic acids encoding the ΤI

The present invention is directed to novel polypeptides and to nucleic ABacid molecules encoding those polypeptides. Also provided herein are vectors and host cells comprising those nucleic acid sequences, chimeric polypeptide molecules comprising the polypeptides of the present invention fused to heterologous polypeptide sequences, antibodies which bind to the polypeptides of the present invention and to methods for producing the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:3495 USPATFULL

TITLE:

Secreted and transmembrane polypeptides and nucleic

acids encoding the same

INVENTOR(S):

Ashkenazi, Avi, San Mateo, CA, UNITED STATES Botstein, David, Belmont, CA, UNITED STATES Desnoyers, Luc, San Francisco, CA, UNITED STATES Eaton, Dan L., San Rafael, CA, UNITED STATES

Ferrara, Napoleone, San Francisco, CA, UNITED STATES Filvaroff, Ellen, San Francisco, CA, UNITED STATES

Fong, Sherman, Alameda, CA, UNITED STATES Gao, Wei-Qiang, Palo Alto, CA, UNITED STATES

Gerber, Hanspeter, San Francisco, CA, UNITED STATES Gerritsen, Mary E., San Mateo, CA, UNITED STATES Goddard, Audrey, San Francisco, CA, UNITED STATES Godowski, Paul J., Burlingame, CA, UNITED STATES Grimaldi, J. Christopher, San Francisco, CA, UNITED

**STATES** 

Gurney, Austin L., Belmont, CA, UNITED STATES

Hillan, Kenneth J., San Francisco, CA, UNITED STATES Kljavin, Ivar J., Lafayette, CA, UNITED STATES Mather, Jennie P., Millbrae, CA, UNITED STATES

Pan, James, Belmont, CA, UNITED STATES

Paoni, Nicholas F., Belmont, CA, UNITED STATES Roy, Margaret Ann, San Francisco, CA, UNITED STATES Stewart, Timothy A., San Francisco, CA, UNITED STATES

Tumas, Daniel, Orinda, CA, UNITED STATES

Williams, P. Mickey, Half Moon Bay, CA, UNITED STATES Wood, William I., Hillsborough, CA, UNITED STATES

Genentech, Inc. (U.S. corporation) PATENT ASSIGNEE(S):

> NUMBER KIND DATE

PATENT INFORMATION: APPLICATION INFO.:

US 2003003530 A1 20030102 US 2001-904011 A1 20010711 20010711 (9)

RELATED APPLN. INFO.: Continuation of Ser. No. US 2000-665350, filed on 18

	T
אירו ארטו אם	TNFORMATION ·

	NUMBER	DATE	
		10000010	
WO	1998-US18824 1998-US19177		
WO	1998-US19377	19980914 19980916	
WO	1998-US19437	19980917	
WO	1998-US25108	19981201	
WO	1999-US20594	19990908	
WO	1999-US20944	19990913	
WO	1999-US21090	19990915	
WO	1999-US21547	19990915	
WO	1999-US23089	19991005	
WO	1999-US28214 1999-US28313	19991129 19991130	
WO	1999-US28301	19991201	
WO	1999-US28564	19991202	
WO	1999-US28565	19991202	
WO	1999-US30095	19991216	
WO		19991220	
WO		19991220	
WO	2000-US219	20000105	
WO	2000-US3565	20000211	
WO	2000-US4414 2000-US5004	20000222	
WO	2000-055004 2000-US5841	20000224	
WO	2000 US3341 2000-US7377	20000320	
WO	2000-US8439	20000330	
WO	2000-US14042	20000522	
WO	2000-US15264	20000602	
WO	2000-US20710	20000728	
MO	2000-US23328	20000824	1001
US US	1997-59115P 1997-59184P	19970917 19970917	(60) (60)
US	1997-59184P 1997-59122P	19970917	(60)
US	1997-59117P	19970917	(60)
US	1997-59113P	19970917	(60)
US	1997-59121P	19970917	(60)
US		19970917	
	1997-59263P	19970918	
US		19970918	(60)
US US	1997-62125P 1997-62287P	19971015 19971017	(60) (60)
US	1997-62285P	19971017	(60)
US	1997-63486P	19971021	(60)
US	1997-62816P	19971024	(60)
US	1997-62814P	19971024	(60)
US	1997-63127P	19971024	(60)
US	1997-63120P	19971024	(60)
US	1997-63121P	19971024	(60)
US US	1997-63045P 1997-63128P	19971024 19971024	(60) (60)
US	1997-63128P	19971024	(60)
US	1997-63327P	19971027	(60)
US	1997-63549P	19971028	(60)
US	1997-63541P	19971028	(60)
US	1997-63550P	19971028	(60)
US	1997-63542P	19971028	(60)
US	1997-63544P	19971028	(60)
US	1997-63564P	19971028	(60)
US US	1997-63734P 1997-63738P	19971029 19971029	(60) (60)
US	1997-63704P	19971029	(60)

19971029 (60) US 1997-63435P US 1997-64215P 19971029 (60) US 1997-63735P 19971029 (60) US 1997-63732P 19971029 (60) US 1997-64103P 19971031 (60) US 1997-63870P 19971031 (60) US 1997-64248P 19971103 (60) US 1997-64809P 19971107 (60) US 1997-65186P 19971112 (60) US 1997-65846P 19971117 (60) US 1997-65693P 19971118 (60) 19971121 (60) US 1997-66120P 19971121 (60) US 1997-66364P 19971124 (60) US 1997-66772P US 1997-66466P 19971124 (60) 19971124 (60) US 1997-66770P US 1997-66511P 19971124 (60) US 1997-66453P 19971124 (60)

DOCUMENT TYPE:

Utility APPLICATION

FILE SEGMENT: LEGAL REPRESENTATIVE:

BRINKS HOFER GILSON & LIONE, P.O. BOX 10395, CHICAGO,

IL, 60610

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 38

NUMBER OF DRAWINGS:

124 Drawing Page(s)

LINE COUNT:

21255

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L5 ANSWER 10 OF 76 USPATFULL

TI Secreted and transmembrane polypeptides and nucleic acids encoding the

The present invention is directed to novel polypeptides and to nucleic acid molecules encoding those polypeptides. Also provided herein are vectors and host cells comprising those nucleic acid sequences, chimeric polypeptide molecules comprising the polypeptides of the present invention fused to heterologous polypeptide sequences, antibodies which bind to the polypeptides of the present invention and to methods for producing the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2002:344632 USPATFULL

TITLE:

Secreted and transmembrane polypeptides and nucleic

acids encoding the same

INVENTOR (S):

Ashkenazi, Avi, San Mateo, CA, UNITED STATES
Botstein, David, Belmont, CA, UNITED STATES
Desnoyers, Luc, San Francisco, CA, UNITED STATES
Eaton, Dan L., San Rafael, CA, UNITED STATES
Ferrara, Napoleone, San Francisco, CA, UNITED STATES
Filvaroff, Ellen, San Francisco, CA, UNITED STATES
Fong, Sherman, Alameda, CA, UNITED STATES
Gao, Wei-Qiang, Palo Alto, CA, UNITED STATES
Gerber, Hanspeter, San Francisco, CA, UNITED STATES
Gerritsen, Mary E., San Mateo, CA, UNITED STATES
Goddard, Audrey, San Francisco, CA, UNITED STATES
Godowski, Paul J., Burlingame, CA, UNITED STATES
Grimaldi, J. Christopher, San Francisco, CA, UNITED
STATES

Gurney, Austin L., Belmont, CA, UNITED STATES Hillan, Kenneth J., San Francisco, CA, UNITED STATES Kljavin, Ivar J., Lafayette, CA, UNITED STATES Mather, Jennie P., Millbrae, CA, UNITED STATES Pan, James, Belmont, CA, UNITED STATES

Paoni, Nicholas F., Belmont, CA, UNITED STATES Roy, Margaret Ann, San Francisco, CA, UNITED STATES Stewart, Timothy A., San Francisco, CA, UNITED STATES Tumas, Daniel, Orinda, CA, UNITED STATES Williams, P. Mickey, Half Moon Bay, CA, UNITED STATES Wood, William I., Hillsborough, CA, UNITED STATES

PATENT ASSIGNEE(S):

Genentech, Inc. (U.S. corporation)

	NUMBER	KIND	DATE	
PATENT INFORMATION: APPLICATION INFO.:	US 2002198366 US 2001-907841	A1 A1	20021226 20010717	

APPLICATION INFO.: US 2001-907841 Al 20010717 (9)
RELATED APPLN. INFO.: Continuation of Ser. No. US 2000-665350, filed on 18

Sep 2000, PENDING

# PRIORITY INFORMATION:

NUMBER	DATE
	10000010
WO 1998-US18824 WO 1998-US19177	19980910 19980914
WO 1998-US19330	19980916
WO 1998-US19437	19980917
WO 1998-US25108	19981201
WO 1999-US20594	19990908
WO 1999-US20944	19990913
WO 1999-US21090	19990915
WO 1999-US21547	19990915
WO 1999-US23089 WO 1999-US28214	19991005 19991129
WO 1999-US28313	19991130
WO 1999-US28301	19991201
WO 1999-US28564	19991202
WO 1999-US28565	19991202
WO 1999-US30095	19991216
WO 1999-US30999	19991220
WO 1999-US30911	19991220
WO 2000-US219	20000105 20000211
WO 2000-US3565 WO 2000-US4414	20000211
WO 2000-US5004	20000222
WO 2000-US5841	20000402
WO 2000-US7377	20000320
WO 2000-US8439	20000330
WO 2000-US14042	20000522
WO 2000-US15264	20000602
WO 2000-US20710	20000728
WO 2000-US23328 US 1997-59115P	20000824 19970917 (60)
US 1997-59184P	19970917 (60)
US 1997-59122P	19970917 (60)
US 1997-59117P	19970917 (60)
US 1997-59113P	19970917 (60)
US 1997-59121P	19970917 (60)
US 1997-59119P	19970917 (60)
US 1997-59263P	19970918 (60)
US 1997-59266P US 1997-62125P	19970918 (60) 19971015 (60)
US 1997-62125P	19971013 (60)
US 1997-62285P	19971017 (60)
US 1997-63486P	19971021 (60)
US 1997-62816P	19971024 (60)
US 1997-62814P	19971024 (60)
US 1997-63127P	19971024 (60)
US 1997-63120P	19971024 (60)
US 1997-63121P	19971024 (60)
US 1997-63045P US 1997-63128P	19971024 (60) 19971024 (60)
03 1337-031202	133/1024 (00)



.....

# WEST

1

Generate Collection

L2: Entry 3 of 5

File: USPT

Feb 3, 1998

DOCUMENT-IDENTIFIER: US 5713374 A TITLE: Fixation method for the attachment of wound repair materials to cartilage defects

BSPR:

Techniques were developed to utilize autologous tissue, such as transplantation of: 1) osteochondral grafts (DePalma, et al. 1963); 2) chondrocytes (Grande, et al. 1989); 3) periosteum (Homminga, et al., 1990); and 4) demineralized bone (Dahlberg and Kreicbers, 1991). These techniques have been used to transplant whole or partial joints, with mixed results. For example, a number of investigators attempted to heal cartilage defects using chondrocytes isolated from epiphysial plates, as well as articular cells, with the hypothesis that these cells would have a greater chance of success due to their heightened metabolism (Itay, et al. 1987). Clinical studies using cultured cells reported excellent results, showing a significant decrease in pain and restoration of normal function after two to four years post-op (Iloika, et al. 1990; Ilomminga, et al. 1990).

# **End of Result Set**

Generate Collection

L2: Entry 5 of 5

File: USPT

Feb 10, 1987

DOCUMENT-IDENTIFIER: US 4642120 A TITLE: Repair of cartilage and bones

#### BSPR:

When articular cartilage is damaged by trauma, infection or degenerative processes, such damages generally fail to heal or even improve. Hitherto various attempts have been made to resort to osteochondral grafts and to the provision of various forms of prosthesis, but long term results have been poor and discouraging. There have been reported attempts to use cultured chondrocytes as a source of cartilage transplants, but integration of the transplants with the neighboring cartilage was generally unsatisfactory.

5 22 6914

Page 1

L16 ANSWER 7 OF 8 MEDLINE

DUPLICATE 4

ACCESSION NUMBER:

97270218

DOCUMENT NUMBER:

CORPORATE SOURCE:

97270218

TITLE:

Regeneration of articular

cartilage defects in rabbits by osteogenic

protein-1 (bone morphogenetic

MEDLINE

protein-7).

AUTHOR:

Grgic M; Jelic M; Basic V; Basic N; Pecina M; Vukicevic S Drago Perovic Institute of Anatomy, School of Medicine,

University of Zagreb, Croatia.

SOURCE:

ACTA MEDICA CROATICA, (1997) 51 (1) 23-7.

Journal code: BH2. ISSN: 1330-0164.

PUB. COUNTRY:

Croatia

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

ENTRY MONTH:

199707

Osteogenic protein-1 (OP-1, BMP-7), a member of the transforming AB growth factor-beta family, induces cartilage and bone formation when implanted at intra and extraskeletal sites in vivo. The human OP-1 gene has been cloned and biologically active recombinant OP-1 homodimers have been produced. In the present study, the authors investigated the influence of OP-1 on healing of full-thickness articular cartilage defects, made by drilling two adjacent (phi 3mm) holes through articular cartilage of NZW rabbit knee joint were dissected and examined histomorphometrically. Results indicated that OP-1 induced articular cartilage healing and regeneration of the joint surface which contained cells resembling mature joint chondrocytes. These data imply a new strategy for biological repair of damaged joint surfaces in humans.

L17 ANSWER 2 OF 4 MEDLINE

ACCESSION NUMBER:

MEDLINE 93101749

DOCUMENT NUMBER:

93101749

TITLE:

Reconstruction of the bone--bone marrow organ by osteogenin, a bone morphogenetic protein, and

demineralized

bone matrix in calvarial defects of adult primates.

**AUTHOR:** 

Ripamonti U; Ma S S; Cunningham N S; Yeates L; Reddi A H Medical Research Council/University of the Witwatersrand,

CORPORATE SOURCE:

Johannesburg, South Africa.

SOURCE:

PLASTIC AND RECONSTRUCTIVE SURGERY, (1993 Jan) 91 (1)

27-36.

Journal code: P9S. ISSN: 0032-1052.

PUB. COUNTRY:

United States Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

Abridged Index Medicus Journals; Priority Journals FILE SEGMENT:

ENTRY MONTH: 199303

Information concerning the efficacy of osteogenin, a bone morphogenetic protein, and demineralized bone matrix in

orthotopic sites in nonhuman primates is a prerequisite for potential clinical application in humans. After exposure of the calvaria, 84

cranial

defects, 25 mm in diameter, were prepared in 26 adult male baboons (Papio ursinus). Defects were implanted with insoluble collagenous bone matrix (ICBM, the inactive collagenous residue after dissociative extraction of bone matrix with 4 M guanidine hydrochloride) reconstituted with osteogenin fractions isolated from baboon bone matrix by chromatography

on

heparin-Sepharose and hydroxyapatite-Ultrogel (Og Hep-HA) or osteogenin further purified using Sephacryl S-200 gel filtration chromatography (Og S-200). Baboon osteogenin with the highest biologic activity in a rodent bioassay, as determined by alkaline phosphatase activity, calcium

content,

and histologic analysis, was used for orthotopic implantation in baboons. Additional defects were implanted with baboon demineralized bone matrix (DBM) or ICBM without osteogenin as control. Defects also were grafted with corticocancellous bone harvested from the iliac crest or left ungrafted to monitor the spontaneous regeneration potential of the adult baboon calvaria. Undecalcified bone sections at 7 microns were prepared from the harvested specimens 30 and 90 days after surgery. Histomorphometry demonstrated that Og S-200 induced copious amounts of bone and osteoid as early as day 30 (P < 0.01 versus ICBM, autogenous grafts and untreated defects). At day 90, in implants of Og S-200, Og Hep-HA, and DBM, bone and marrow formation was extensive, culminating in complete regeneration of the craniotomies. In implants of DBM, bone formed with an intervening phase of cartilage development. This provides the phenotypic evidence of endochondral bone differentiation by induction in defects of membranous calvarial bone in adult primates. These results establish the potential therapeutic application of osteogenin and demineralized bone matrix for the architectural reconstruction of the bone-bone marrow organ in humans.

Page 1

L17 ANSWER 4 OF 4 MEDLINE

ACCESSION NUMBER:

86278360

86278360

DOCUMENT NUMBER: TITLE:

Bone repair induced by bone morphogenetic protein in ulnar

God I

defects in dogs.

AUTHOR:

Nilsson O S; Urist M R; Dawson E G; Schmalzried T P;

Finerman G A

SOURCE:

JOURNAL OF BONE AND JOINT SURGERY. BRITISH VOLUME, (1986

Aug) 68 (4) 635-42.

Journal code: HK7. ISSN: 0301-620X.

PUB. COUNTRY:

ENGLAND: United Kingdom

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Abridged Index Medicus Journals; Priority Journals

ENTRY MONTH: 198611

AB In dogs, resection of a length of the ulna equal to twice the diameter of the mid-shaft leaves a defect which consistently fails to unite. In

response to an implant of 100 mg of bovine bone

morphogenetic protein (BMP), the defect

becomes filled by callus consisting of fibrocartilage, cartilage and woven bone within four weeks. The cartilage is resorbed and replaced by new bone in four to eight weeks. Woven bone is then resorbed, colonised by bone marrow cells and remodelled into lamellar bone. Union

of

the defect is produced by 12 weeks. Control defects filled with autogeneic

cortical bone chips unite after the same period. In regeneration induced by bone morphogenetic protein (

BMP) and in repair enhanced by bone graft, union depends upon the proliferation of cells within and around the bone ends. Our working hypothesis is that BMP induces the differentiation of perivascular connective tissue cells into chondroblasts and osteoprogenitor cells and thereby augments the process of bone regeneration from the cells already present in the endosteum and periosteum.

(20) 1/27

DUPLICATE 1

Late

L17 ANSWER 1 OF 4 MEDLINE

96023917

96023917

MEDLINE

DOCUMENT NUMBER:

**:** 

TITLE:

Commercially-prepared allograft material has biological

activity in vitro.

AUTHOR:

Shigeyama Y; D'Errico J A; Stone R; Somerman M J Department of Periodontics/Prevention/Geriatrics,

University of Michigan, Ann Arbor, USA.

CONTRACT NUMBER:

CORPORATE SOURCE:

ACCESSION NUMBER:

DE09532 (NIDCR)

SOURCE:

JOURNAL OF PERIODONTOLOGY, (1995 Jun) 66 (6) 478-87.

Journal code: JMT. ISSN: 0022-3492.

PUB. COUNTRY:

United States

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals; Dental Journals

ENTRY MONTH: 199601

AB The well-established finding that implantation of demineralized bone matrix at non-skeletal sites results in formation of cartilage and bone has been attributed to bone morphogenetic proteins/factors. Commercially-available demineralized bone allograft materials are being used currently to reconstruct/regenerate bone. The studies described here focused on establishing biological activity of protein extracts prepared from commercially obtained bone graft material in vitro. Furthermore, the biological activity of these protein extracts in vitro was compared with similar extracts prepared from freshly obtained

human bone. Biological activities of bone matrix proteins examined included their ability to promote proliferation, attachment, and migration

of gingival fibroblasts using an in vitro system. Guanidine followed by quanidine/EDTA was used to separate bone matrix proteins into proteins associated with soft tissues of bone and proteins retained within the mineral compartment, respectively. Two preparations of each starting material were tested and the biological activity of each preparation was evaluated in triplicate at least three times. Slot blot analysis revealed that commercially-prepared material contained type I collagen; fibronectin; BSP; and BMP-2, 4, and 7. However, the freshly prepared bone extracts appeared to have higher BMP concentrations. The ability of commercial extracts to promote cell proliferation, while significant, was limited and significantly less when compared with similar extracts prepared from freshly obtained bone. All extracts promoted cell attachment significantly, while none of the extracts promoted cell migration. Thus, commercially-prepared material retained proteins having the capacity to influence cell behavior in vivo. However, some biological activity as measured in vitro was lost as a result of tissue processing.

Pt. 36

Page 1

DUPLICATE 2

MEDLINE MEDINE

L16 ANSWER 5 OF 8 MEDLINE

ACCESSION NUMBER:

1998258985

DOCUMENT NUMBER:

98258985

TITLE: Osteogen

Osteogenic protein (OP-1, BMP-7) stimulates cartilage differentiation of human and goat perichondrium tissue in

vitro.

AUTHOR:

Klein-Nulend J; Louwerse R T; Heyligers I C; Wuisman P I;

Semeins C M; Goei S W; Burger E H

CORPORATE SOURCE:

ACTA-Vrije Universiteit, Department of Oral Cell Biology,

Amsterdam, The Netherlands..

J.Klein\_Nulend.OCB.ACTA@med.vu

.nl

SOURCE:

JOURNAL OF BIOMEDICAL MATERIALS RESEARCH, (1998 Jun 15) 40

(4) 614-20.

Journal code: HJJ. ISSN: 0021-9304.

PUB. COUNTRY:

United States

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

199811

ENTRY WEEK:

19981104

AB The objective of this study was to examine in vitro the influence of recombinant human osteogenic protein-1 [rhOP-1, or bone

morphogenetic protein-7 (BMP-7)] on cartilage

formation by human and goat perichondrium tissue containing progenitor cells with chondrogenic potential. Fragments of outer ear perichondrium tissue were embedded in clotting autologous blood to which rhOP-1 had

been

added or not added (controls), and the resulting explant was cultured for 3 weeks without further addition of rhOP-1. Cartilage formation was monitored biochemically by measuring [35S]-sulphate incorporation into proteoglycans and histologically by monitoring the presence of metachromatic matrix with cells in nests. The presence of rhOP-1 in the explant at the beginning of culture stimulated [35S]-sulphate incorporation into proteoglycans in a dose-dependent manner after 3 weeks of culture. Maximal stimulation was reached at 40 microg/mL (human explants: +148%; goat explants: +116%). Histology revealed that explants treated with 20-200 microg/mL of rhOP-1, but not untreated control explants, contained areas of metachromatic-staining matrix with chondrocytes in cell nests. It was concluded that rhOP-1 stimulates differentiation of cartilage from perichondrium tissue. The direct

actions

of rhOP-1 on perichondrium cells in the stimulation of chondrocytic differentiation and production of cartilage matrix in vitro provides a cellular mechanism for the induction of cartilage formation by rhOP-1 in vivo. Thus rhOP-1 may promote early steps in the cascade of events

leading

to cartilage formation and could prove to be an interesting factor in the regeneration of cartilage in articular cartilage defects.

Mich 10 20